

Sa247851_0026.Dna
Gcg_Geneseq_D:T19113

XP 2128439

ID T19113 standard; cDNA to mRNA; 351 BP.
AC T19113;
DT 04-JUL-1996 (first entry)

DE Human gene signature HUMGS00127.
KW Gene signature; messenger RNA; mRNA; relative abundance; frequency;
KW human; cloning; mapping; non-biased library; diagnosis; detection;
KW cell typing; abnormal cell function; ss.
OS Homo sapiens.
PN WO 9514772-A1.
PD 01-JUN-1995.
PF 11-NOV-1994; 94WO-JP01916.
PR 12-NOV-1993; 93JP-0355504.
PA (MATS/) MATSUBARA K.
PA (OKUB/) OKUBO K.
PI Matsubara K, Okubo K;
DR WPI; 1995-206931/27.
PT Identifying gene signatures in 3'-directed human cDNA library - e.g.
PT for diagnosis of abnormal cell function, by preparing cDNA that
PT reflects relative abundance of corresp. mRNA in specific human

PT tissues
PS Claim 1; Page 304; 2245pp; Japanese.
CC A single-stranded DNA (or its complementary strand or the corresp.
CC double-stranded DNA) which comprises one of the 7837 "GS" sequences
CC given in T19001-T26837 and which is able to hybridise to part of
CC human genomic DNA, cDNA or mRNA is claimed. The GS (Gene Signature)
CC sequences were obtained from 3'-directed cDNA libraries prepared
CC from various human tissues; synthesis of cDNA was initiated from the
CC 3'-end of mRNA by using poly(T) as the sole primer. Since the 3'-
CC untranslated sequence is unique to a particular mRNA species, almost
CC all the 3'-oriented cDNAs hybridise with specific mRNAs. Each library
CC is constructed so as to reflect accurately the relative abundance of
CC different mRNAs in the particular tissue from which it was derived.
CC The appearance frequency of a given GS in a cDNA library can be
CC determined (esp. using primers and probes derived from the GS
CC sequences) as a means of diagnosing abnormal cell function or for
CC recognising different cell types.
SQ Sequence 351 BP; 123 A; 45 C; 58 G; 121 T; 4 other;

SCORES Init1: 1638 Initn: 1638 Opt: 1706 z-score: 597.6 E(): 5.9e-24
98.9% identity in 349 bp overlap

1480 1490 1500 1510 1520 1530
Sa247851_002 TTATCTGCCATTAAAAGTATCCAGTATTGATCACATTATAAAT-AATGAAAAAAAT
||| ||| ||| ||| ||| |||

T19113 GATCACATTATNATAAATAATGAAAAAAAT
10 20 30

1540 1550 1560 1570 1580 1590
Sa247851_002 GATTTAATCTGTAATAAACCTGGTTATTGTGCAGTGACTGTAATATACTAGAGTTATAAT
||| ||| ||| ||| ||| |||

T19113 GATTTAATCTGTAATAAACCTGGTTATTGTGCAGTGACTGTAATATACTAGAGTTATAAT
||| ||| ||| ||| |||

40 50 60 70 80 90

1600 1610 1620 1630 1640 1650
Sa247851_002 AAATTGTTACTCTGCCTCACCAAACACATGCTAGGATATAACCCCCAAAATAAGTATT
T19113 AAATTGTTACTCTGCCTCACCAAACACATGCTAGGATATAACCCCCAAAATAAGTATT
100 110 120 130 140 150

1660 1670 1680 1690 1700 1710
Sa247851_002 AACTTGCATTAGGTATAAAGGAGACTGGGTGCTATAATTAGATTATTTGAGGCAGACA
T19113 AACCTTGCAATTAGGTATAAAGGAGACTGGGTGCTATAATNAGATTATTTGAGGCAGACA
160 170 180 190 200 210

1720 1730 1740 1750 1760 1770
Sa247851_002 GAGAGCTGTTATCCTAAC TGATTAGTATGTTCTGTAAATTGAGAAAATGTTACCAAATT
T19113 GAGAGCTGTTATCCTAAC TGATTAGTATGTTCTGTAAATTGAGAAAATGTTACCAAATT
220 230 240 250 260 270

1780 1790 1800 1810 1820 1830
Sa247851_002 ATACTTTTAGTGATTACATGTACATTTATAGGGGACATGTTCTGTGTATAGCGAATA
T19113 ATACTTTTAGTGATTACATGTACATTTATAGGGGACATGTTCTGTGTATAGCGAATA
280 290 300 310 320 330

1840 1850 1860 1870
Sa247851_002 AATAACTTTATAGTATCAAAAAAAAAAAAAAA
T19113 AATAACTTTATAGTATCACN
340 350



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(71) Applicant: **Matsubara, Kenichi**
Room 804, 18-1, Yamadaishi-gashi 3-chome
Sulta-shi,
Osaka 565 (JP)
Applicant: **Okubo, Kousaku**
11-26, Segawa 2-chome
Minoo-shi,
Osaka 562 (JP)

(72) Inventor: **Matsubara, Kenichi**
Room 804, 18-1, Yamadaishi-gashi 3-chome
Sulta-shi,
Osaka 565 (JP)
Inventor: **Okubo, Kousaku**
11-26, Segawa 2-chome
Minoo-shi,
Osaka 562 (JP)

(74) Representative: **Vossius, Tilman et al**
Dr. Volker Vossius,
Patent- und Rechtsanwaltskanzlei,
Holbeinstrasse 5
D-81679 München (DE)

(54) GENE SIGNATURE.

(57) A 3'-directed cDNA library which accurately reflects the abundance ration of mRNA in a cell has been prepared from various human tissues, and sequencing of the cDNAs contained in the library has be conducted to examine the incidence of each cDNA in each tissue. As each cDNA has expression information with each tissue corresponding to the mRNA concentration, these cDNAs are usable as a probe or primer for detecting cell anomoly or discriminating cells. The cloned gene can produce proteins utilizable as a medicine or the like.

Table 1

SEQUENCE LISTING

(1) GENERAL INFORMATION:

5 (i) APPLICANT:

- (A) NAME: CHUGAI PHARMACEUTICAL CO., LTD.
- (B) STREET: 41-8, Takada 3-chrome, Toshima-ku
- (C) CITY: Tokyo
- 10 (E) COUNTRY: JAPAN
- (F) ZIP: 171

(ii) TITLE OF INVENTION: GENE SIGNATURE

15 (iii) NUMBER OF SEQUENCES: 7848

(iv) COMPUTER READABLE FORM:

- (A) MEDIUM TYPE: Diskette, 3.5 in., DS, 1.44 MB
- (B) COMPUTER: IBM PC compatible
- 20 (C) OPERATING SYSTEM: PC-DOS/ MS-DOS
- (D) SOFTWARE: MS-DOS

(v) CURRENT APPLICATION DATA

- (A) APPLICATION NUMBER: EP 95900295.7

25 (vi) PRIOR APPLICATION DATA

- (A) APPLICATION NUMBER: PCT/JP94/01916
- (B) FILING DATE: 11. November 1994

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SEQ ID NO:7844
 SEQUENCE LENGTH:37
 SEQUENCE TYPE:nucleic acid
 5 STRANDEDNESS:single
 TOPOLOGY:linear
 SEQUENCE DESCRIPTION:
 CTCGCTGCC CATCCTTATA CAGGCTCACT TTTGTCT 37

10 SEQ ID NO:7845
 SEQUENCE LENGTH:37
 SEQUENCE TYPE:nucleic acid
 15 STRANDEDNESS:single
 TOPOLOGY:linear
 SEQUENCE DESCRIPTION:
 CTCGCTGCC CATGTATAGG GACAGCATT CTGAGAG 37

20 SEQ ID NO:7846
 SEQUENCE LENGTH:38
 SEQUENCE TYPE:nucleic acid
 25 STRANDEDNESS:single
 TOPOLOGY:linear
 SEQUENCE DESCRIPTION:
 CTGGTTCGGC CCACCTCTGA AGGTTCCAGA ATCGATAG 38

30 SEQ ID NO:7847
 SEQUENCE LENGTH:22
 SEQUENCE TYPE:nucleic acid
 35 STRANDEDNESS:single
 TOPOLOGY:linear
 SEQUENCE DESCRIPTION:
 CCAGGGTTTT CCCAGTCACG AC 22

40 SEQ ID NO:7848
 SEQUENCE LENGTH:22
 SEQUENCE TYPE:nucleic acid
 45 STRANDEDNESS:single
 TOPOLOGY:linear
 SEQUENCE DESCRIPTION:
 TCACACAGGA AACAGCTATG AC 22

50 Claims

1. A purified single-stranded DNA, a purified single-stranded DNA complementary thereto, or a purified double-stranded DNA consisting of said single strands, containing all or a portion of a single-stranded DNA or a single-stranded DNA complementary thereto comprising any of the base sequences listed under SEQ ID NO 1-7837 and hybridizing specifically to a particular site of human genomic DNA, human cDNA or human mRNA.

2. A DNA probe consisting of a purified single-stranded DNA , a purified single-stranded DNA complementary thereto, or a purified double-stranded DNA consisting of said single strands, containing all or a portion of a single-stranded DNA or a single-stranded DNA complementary thereto comprising any of the base sequences listed under SEQ ID NO 1-7837 and hybridizing specifically to a particular site of human genomic DNA, human cDNA or human mRNA.

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3. A DNA primer consisting of a purified single-stranded DNA, a purified single-stranded DNA complementary thereto, or a purified double-stranded DNA consisting of said single strands, containing all or a portion of a single-stranded DNA or a single-stranded DNA complementary thereto comprising any of the base sequences listed under SEQ ID NO 1-7837 and hybridizing specifically to a particular site of human genomic DNA, human cDNA or human mRNA.

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4. A purified single-stranded DNA, a purified single-stranded DNA complementary thereto, or a purified double-stranded DNA consisting of said single strands, containing all or a portion of a single-stranded DNA or a single-stranded DNA complementary thereto, wherein said single-stranded DNA is complementary to a human mRNA containing any of the base sequences listed under SEQ ID NO 1-7837 (wherein T is read as U) or any portion thereof at its 3' region, and hybridizing specifically to a particular site of human genomic DNA, human cDNA or human mRNA.

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5. A DNA probe consisting of a purified single-stranded DNA, a purified single-stranded DNA complementary thereto, or a purified double-stranded DNA consisting of said single strands, containing all or a portion of a single-stranded DNA or a single-stranded DNA complementary thereto, wherein said single-stranded DNA is complementary to a human mRNA containing any of the base sequences listed under SEQ ID NO 1-7837 (wherein T is read as U) or any portion thereof at its 3' region, and hybridizing specifically to a particular site of human genomic DNA, human cDNA or human mRNA.

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6. A DNA primer consisting of a purified single-stranded DNA, a purified single-stranded DNA complementary thereto, or a purified double-stranded DNA consisting of said single strands, containing all or a portion of a single-stranded DNA or a single-stranded DNA complementary thereto, wherein said single-stranded DNA is complementary to a human mRNA containing any of the base sequences listed under SEQ ID NO 1-7837 (wherein T is read as U) or any portion thereof at its 3' region, and hybridizing specifically to a particular site of human genomic DNA, human cDNA or human mRNA.

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Fig. 1

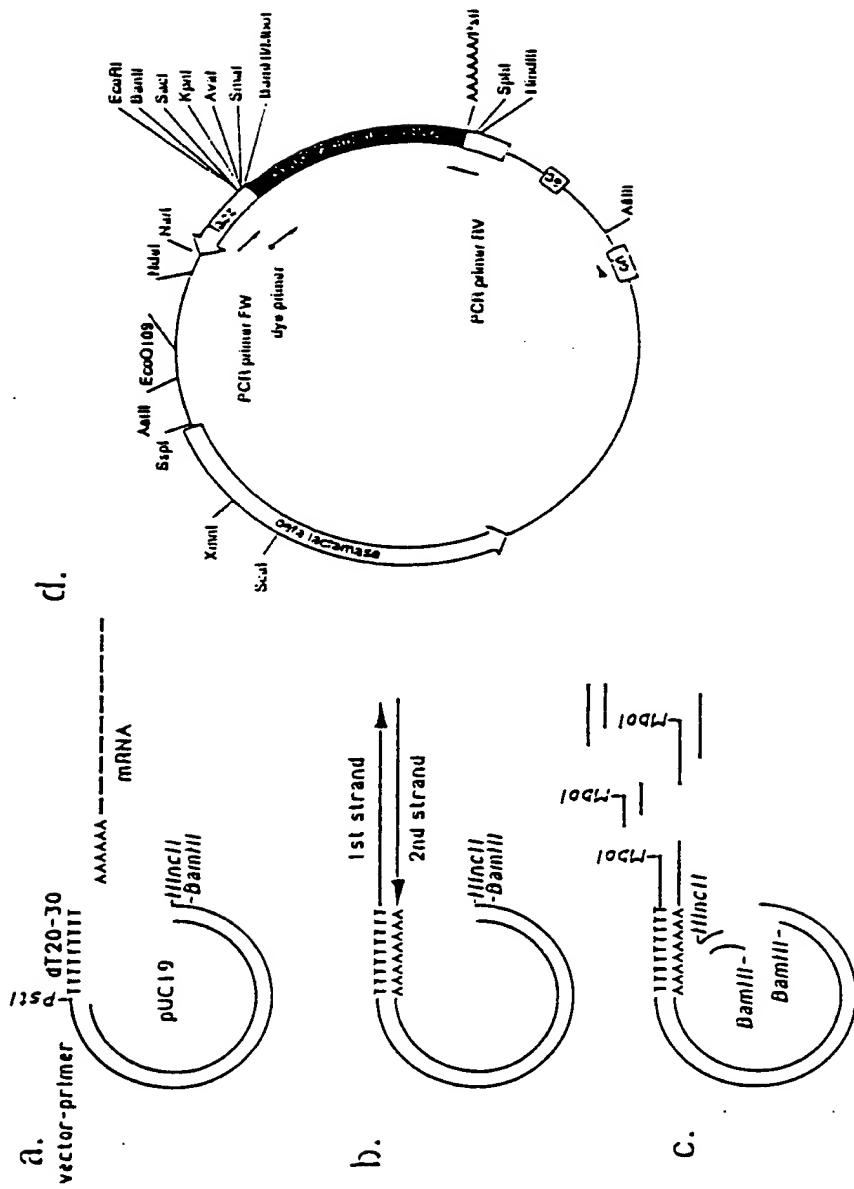


Fig. 12

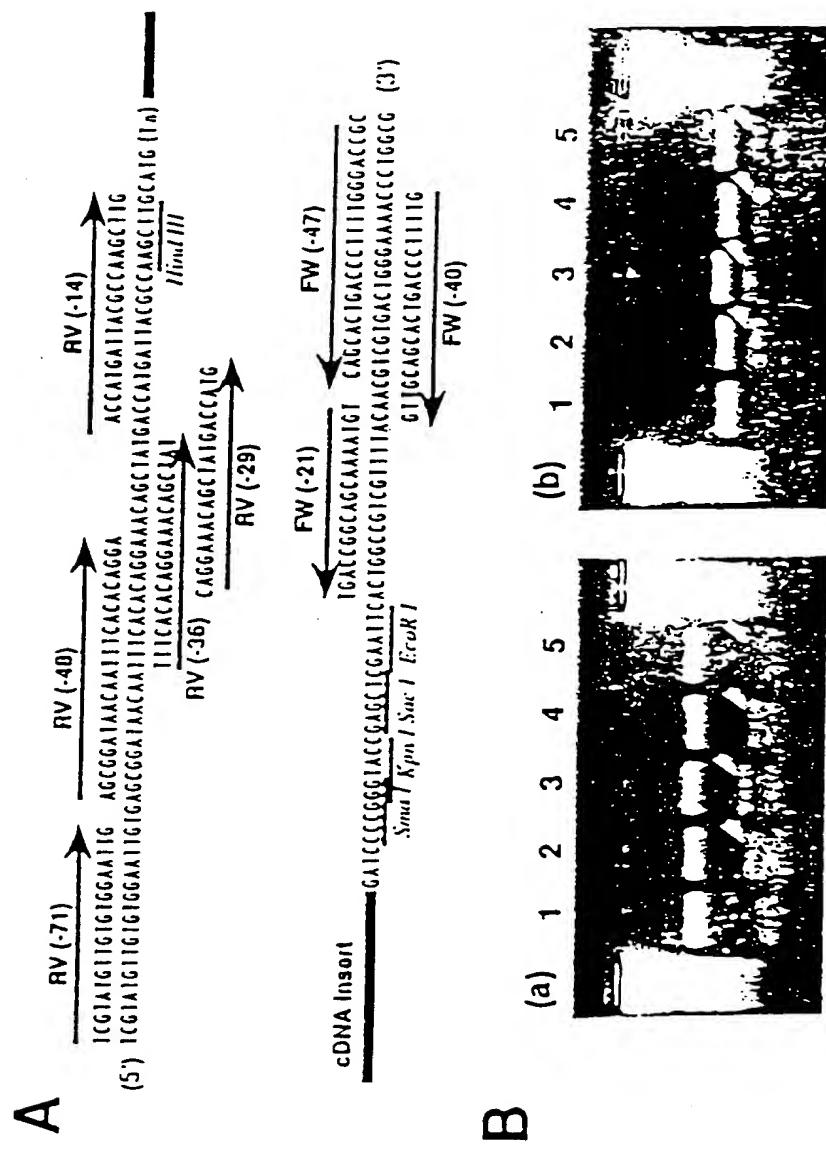


Fig. 3

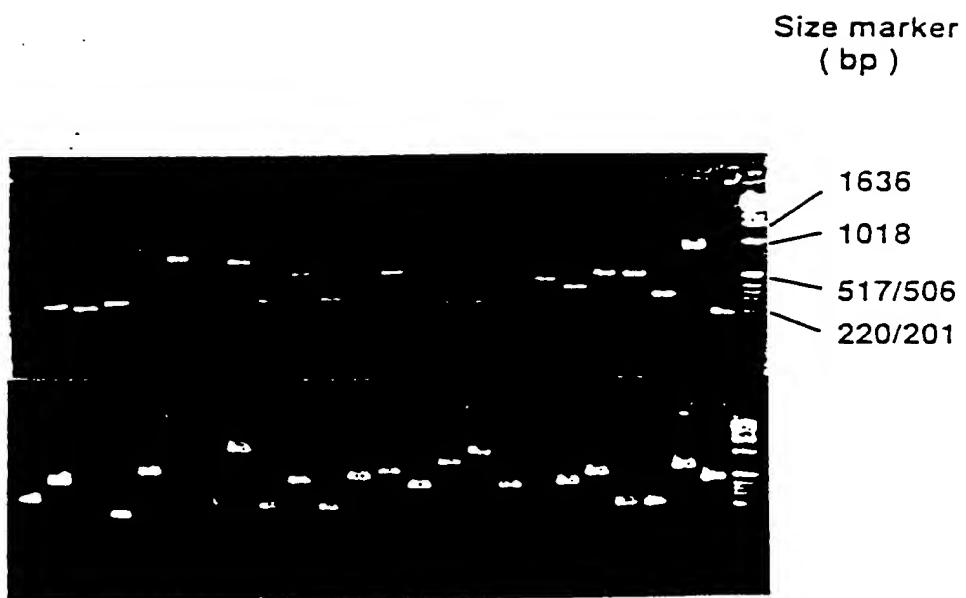


Fig. 4

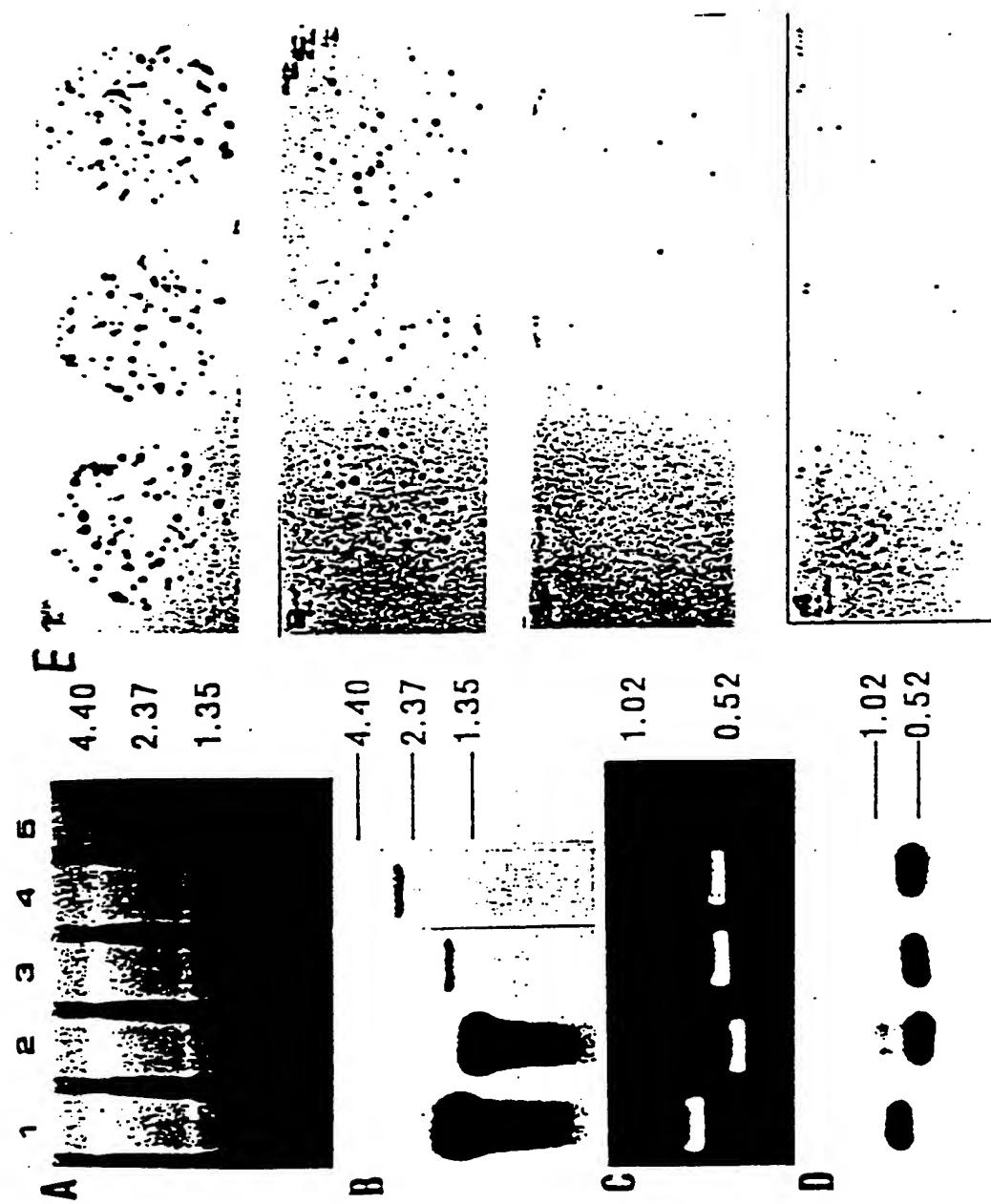


Fig. 4

F

probe No.	1	2	3	4
gene	Elongation factor 1- α	α 1-antitrypsin	HnRNP core protein A1	Inter- α -tryptsin inhibitor
(a) Band intensity of Northern blot(cpm)	687	423	10	15
(b) Band intensity of control blot(cpm)	133	177	100	127
(c) Normalized signal(a)/(b) $\times 10$	52	24	1	1.2
(d) Positive signals on colony blot	307	119	7	9
(e) Relative representation	44	17	1	1.3

Fig. 5

Appearance frequencies of various cDNAs in the 3'-directed HepG2 cDNA library

Group	Clone	Gene	A in 902 (%)	B "in 0,800 (%)"	C "in 26,400 (%)"
	a15	Elongation factor - 1 α	22 (2.2)	307 (3.5)	NT
	c321	Translationally restricted tumor protein	12 (1.2)	89 (1.0)	NT
I	lb030	α -1-antitrypsin	0 (0.0)	119 (1.4)	NT
	hm01b02	Light chain of ferritin	6 (0.6)	62 (0.7)	NT
	c13a04	NADP(II) Menadione oxidoreductase	4 (0.4)	27 (0.3)	NT
	hm02d02	Ribosomal protein S11	3 (0.3)	29 (0.3)	NT
	lb042	Human RNP core protein A1	2 (0.2)	7 (0.1)	NT
	s155	unknown	1	2	5 (0.02)
	s159	unknown	1	2	4 (0.02)
	s639	unknown	1	1	3 (0.01)
	s635	unknown	1	0	2 (0.01)
II	s170	unknown	1	0	1 (0.001)
	s154	unknown	1	0	1 (0.004)
	s167	unknown	1	0	1 (0.004)
	s645	unknown	1	0	1 (0.001)
	s647	unknown	1	0	1 (0.001)
	s632	unknown	0	0 (<0.001)	0 (<0.001)

Fig. 6

GS	CI	Chromosomal position	Sequences of primers		Antisense	AT	TTO	TTC	TAC	G	T
			Sense	Annealing							
61620184	pm12166	-	CAGGCCCGTAACTAT AATGGGACAGTTAACCTA	AAGTTATGTGGGTACAG CCAGCTCCCTGACTGAGA	48	114	115	104	110	1	2
91001024	pm02444	-	TGGACTGGATACCTATC GICACTCTGCCATGAC	ACAGTACCCCTGAAGCT ACCACTCTGCCACACTT	48	81	81	>200	1	-	-
916001035	pm00081	-	GCCCCTAACGAGGAAC GGGTTTCAATGGGTGACC	TAATTCACCTCCGTAC GGCCCACTGIGAAACG	50	104	104	103	107	4	4
91001047	pm11722	-	TTAGGAGCCCTATGGACC TCAGGTCGCTGGAGATG	ATAATCTGGTTAGCATTAC AACATCAGCACAGTAITG	51	114	116	>200	200	6	6
91001054	pm02447	-	AGGTGATGTTACAGGAT CCAAAGCTCCGGTGAA	GGCTGACATCAGCTTGG TACGGGAGGCTATGAA	49	95	95	78	107	-	-
91001116	pm11711	-	TTAGGAGCCCTATGGACC TCAGGTCGCTGGAGATG	ATAATCTGGTTAGCATTAC AACATCAGCACAGTAITG	51	120	122	>200	>200	-	-
91001191	pm06009	-	AGGTGATGTTACAGGAT CCAAAGCTCCGGTGAA	GGCTGACATCAGCTTGG TACGGGAGGCTATGAA	47	97	97	-	200	-	-
91001200	pm11551	-	TTAGGAGCCCTATGGACC TCAGGTCGCTGGAGATG	ATAATCTGGTTAGCATTAC AACATCAGCACAGTAITG	51	120	122	>200	>200	-	-
91001116	pm00082	-	AGGTGATGTTACAGGAT CCAAAGCTCCGGTGAA	GGCTGACATCAGCTTGG TACGGGAGGCTATGAA	47	130	125	>200	>200	-	-
91001146	pm1510	-	TTAGGAGCCCTATGGACC TCAGGTCGCTGGAGATG	ATAATCTGGTTAGCATTAC AACATCAGCACAGTAITG	51	100	100	>200	-	-	-
91001444	pm11349	-	TTAGGAGCCCTATGGACC TCAGGTCGCTGGAGATG	ATAATCTGGTTAGCATTAC AACATCAGCACAGTAITG	40	98	98	-	-	-	-
91001448	pm01277	-	TTAGGAGCCCTATGGACC TCAGGTCGCTGGAGATG	ATAATCTGGTTAGCATTAC AACATCAGCACAGTAITG	43	102,175	100	>200	>200	-	-
91001521	pm11285	-	CCAAATCAAATGTTAAATG CCAGAGGTTCAAGGGATG	GGTACAGATGCAAGACT CCACATGTTGCTTACCTG	46	57	57	78	155	-	-
91001534	pm11291	-	CCAAATCAAATGTTAAATG CCAGAGGTTCAAGGGATG	GGTACAGATGCAAGCT CCACATGTTGCTTACCTG	44	58	58	>200	>200	-	-
91001512	pm12006	1	CCAAATCAAATGTTAAATG CATGATCTCTGGCTGGTA	AAACCTTATGGCTCCTCT AAACAGTAGTGGCAGT	45	84	84	109	-	2	2
61620120	pm11150	2	CCAAATCAAATGTTAAATG AGGTGAAATGGGGCTGCT	CCGGTTATGCTGAACTGCT TICATAATGTTAACAGTA	48	119	119	91	115	-	-
91001046	pm11720	2	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	TAATGATATATGGGGTAA CCAGAAGGAAAGGGAACT	40	90	90	-	-	-	-
91001061	pm00031	2	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	TAATGATATATGGGGTAA CCAGAAGGAAAGGGAACT	45	88	88	-	-	-	-
91001050	pm00025	2	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	TAATGATATATGGGGTAA CCAGAAGGAAAGGGAACT	47	125	125	>200	>200	-	-
91001211	pm12010	2	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	TAATGATATATGGGGTAA CCAGAAGGAAAGGGAACT	43	103	103	-	-	-	-
91001252	pm00035	2	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	TAATGATATATGGGGTAA CCAGAAGGAAAGGGAACT	52	137	137	>200	-	-	-
91001264	pm2003	2	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	TAATGATATATGGGGTAA CCAGAAGGAAAGGGAACT	45	105	108	-	-	-	-
91001264	pm2135	2	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	TAATGATATATGGGGTAA CCAGAAGGAAAGGGAACT	45	67	67	>200	>200	-	-
91001264	pm2135	2	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	TAATGATATATGGGGTAA CCAGAAGGAAAGGGAACT	45	75	75	170	190	-	-
91000875	pm2135	2	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	TAATGATATATGGGGTAA CCAGAAGGAAAGGGAACT	44	68	67	-	-	-	-
91001438	pm11671	2	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	TAATGATATATGGGGTAA CCAGAAGGAAAGGGAACT	43	88	88	-	-	-	-
91001433	pm11245	2	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	TAATGATATATGGGGTAA CCAGAAGGAAAGGGAACT	45	115	115	-	-	-	-
91001535	pm1216	2	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	TAATGATATATGGGGTAA CCAGAAGGAAAGGGAACT	42	90	90	-	-	-	-
91000875	pm00049	3	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	AAAGATTGATTCATACAT GGCCACATGCTTACCTG	44	77	77	-	-	-	-
9100129	pm11727	3	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	AAAGATTGATTCATACAT GGCCACATGCTTACCTG	43	75	75	155	>200	-	-
91001061	pm1022	3	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	AAAGATTGATTCATACAT GGCCACATGCTTACCTG	46	83	83	160	140	-	-
91001214	pm12009	3	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	AAAGATTGATTCATACAT GGCCACATGCTTACCTG	55	105	105	113	>200	-	-
91001219	pm2155	3	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	AAAGATTGATTCATACAT GGCCACATGCTTACCTG	42	120	120	150	>200	-	-
91001252	pm1252	3	AAAGCAATACAAATACCAA TAATGATCAGCTGAAATAG	AAAGATTGATTCATACAT GGCCACATGCTTACCTG	43	90	90	95	95	-	-

Fig. 7

91000448	pm2258	GCCAAAGTTCTCTTGTAT	GTCAGTTTATCAGGCA	42	62	>200	69	2
91001032	pm1151	GTCGAATGACTGTGTTAT	GTCATATTCATCAAA	43	80	-	-	1
91001215	pm0988	AGAAAATTAGCATAGGT	TAGAGTCAGTACCGTG	43	100	110	-	1
91001798	pm2167	ATCAAAGTTTATGCTCA	CATCCCACATACAGTC	43	116	>200	160	1
91000997	pm0904.	TCTGTGAGACAGACAA	TCTAAGGAACAGACAT	43	116	>200	160	1
91000594	pm1889	AAGCATGCCATCACAG	CTAAGGCTGACCCCTCAT	49	101	113	200	1
91001065	pm0119	TCACCAGATTTACAGT	GATACAAAGGGTAAGAT	45	87	>200	>200	1
91001101	pm2164	TTACCTTACGGGCTTAC	AGACAAATCCAAAGC	44	120	-	-	1
91001161	pm1150	ATTGTTGAGGGTTACTA	AGAATGGATGCTTATC	47	89	93	>200	1
91000912	pm2120	ATGTCATAGTCCTCTCA	TCGATCTTCATGCTCT	43	101	99	>200	200
91001226	pm1151	CATGGACAGGACACAG	CTCGGCCCTTCGAGTA	53	102	104	145	200
91001234	pm1216	TAGGAAAACGGAGAGG	AGGAGCTGGGCTGAGGTC	48	65	65	110	>200
91001557	pm1785	TATATGAAATTCAGGCTG	TCTAATACTGCTCCATCT	46	90	90	>200	>200
91001523	pm0285	TTGAACTGTTGCTCAGT	TTAAATACTGCTGTAAT	42	86	70	>200	100
91001523	pm0228	GCACCTAGCCICCAAGT	TTTATACTGCTCAGAGC	49	138	138	>200	200
91001562	pm2169	TCTGATTGCAAGGACCAC	TTTGGATTTATGTCATC	43	62	62	>200	45
91000624	pm0191	GACCTGATGTTGAAATGAT	ACTTACGCTTATGGATT	45	119	119	>200	2
91001145	pm0281	-AGCCAACCTGGGTAATC	CCACGGACAGGTGACTAT	56	139	139	115	>200
91001169	pm0219	ATCATTTGGAGAGCTGA	AGAACACTTATCCACA	45	89	89	130	-
91001279	pm1102	TCAGGAGCTGCTGATTA	TTTGGAGTTAACTGTTTA	44	77	75	170	-
91001287	pm0956	ACAGTATGCGGTGAGCTAG	TCCTTAAATAGCCAGTCAG	47	81	81	105	70
91001176	pm2327	TGACCTCTAATGGGCTCAT	AAACCAAACTGACTAAG	40	99	99	110	180
91001248	pm2108	TGTTATGGATTGGCTCTC	CAAAAGCAAAACGGAGATA	44	95	95	-	85
91000780	pm0995	TTCACATCAAACATACA	CTTGTGTTGTTGIVTCIG	43	55	55	-	-
91001055	pm0559	TTMAGATTCACCCATG	CACATGCTTATGGAAACT	44	74	74	72	73
91001157	pm0347	AGTTATGTCGAAAGTGA	AGAAAAACACTGCTGTGG	45	138	138	>200	200
91001284	pm2245	TGTAAATGCTATCTCTCT	GCACATGCTTCATGCTGT	47	100	100	>200	>200
91000228	pm2661	ATCAAACACAACTCAGA	ACIATATATACGCTCACT	42	117	121	124	95
91001159	pm0180	GAATAGCTGGAGATTTCAC	GGAGATCATACCTCAGA	46	100	100	84	95
91001215	pm0445	AAAGTACCTGATGGACAGCTG	50	153	153	>200	160	-
91001352	pm2031	AGGTGAGGGTTTTCAGC	CACATCACTGGTAGACTA	47	93	93	-	2
91001469	pm0559	ACCCCTCTAGTAAGCTATG	TTATAACCAAATCCAGTA	37	47	47	125	53
91001570	pm2810	CTGAAAGGTTGGAAATTA	TTCTATTTCTTACAGTTTAT	42	75,92	75	145	>200
91000279	pm0266	AGTGTATGAAACCTCTG	GTCTATGAAACGGTGTAGC	48	130	130	>200	1
91001163	pm2156	TCTCCATATCACACAGT	ATGATTCGTAGATGAGCA	49	88	83	>200	120
91001193	pm1193	CACAGCATAAAGATATA	ACCTTAATTTAGTTCTC	46	100	100	-	2
91001215	pm2790	CATCATGTTACAGTCGAG	CGTTTGTCAAAGTATG	44	83	82	93	87
91001274	pm1355	AGATGCGATATCCTCTAG	GGAAACGCAAGAAAGCAACAC	47	87	87	>200	>200
91001203	pm0368	CCAAAGCTGGTACGGTTCAG	TTCATATGAACTTGGGTAC	47	95,165	95	>200	>200
91000153	pm2645	CTAAGATTTAATGCGATTC	AGTATGTTAATGCGATGIA	46	104	104	>200	-

Fig. 8

Fig. 9

91000999		X	CTACCATGTTACCTGATT	TACCCACCACTATTAGCA
91001149	pml1759	X	GGAGGGGAGATAAGTGT	AAAATACTAGAGACTGA
91001161	pml2180	X	TCTTAAGTGACCACTT	GGAGATGAGAACTACAT
91001165	pml5048	X	TAAGCCAGTGAATGCGTAA	GTAAGGTTATCTGCATAGA
91001169	pml1394	X	ATCGCTGAAATACCTG	GGGGAGAGACACATGAC
91001176	pml2249	1,18	GATCCATGGAGTAAAT	AATACAAAGCTAACCCAA
91001181	pml0113	1,2,12,13,Y	TIGGATGACATTCTAT	TITATGTAACAAEACT
91001191	pml2222	1,2,3,5,8,12,14,17,X	TATCAAGCTGAAATGTCAC	TACGAAICCACCCAA
91002003	pml0314	1,26,X	1,3,4,5,9,16	AGTIGAGACGGGGAATG
91000210	pml1461	2,20,21,22	TCCAAATGAGGGGTTA	TITTTATGTTGCTCAAGI
91001154	pml1561	2,4,5,10,12,15,17,20,22,Y	GACCTGTTGACATCTGACT	TTAAIGGTTGTTACCTGG
91000216	pml2295	20,X	GCCTGTTATTTCACCTC	ATCTCCCTTGCCTCAGTA
91001077	pml0943	2,5,14,C	TCTGAGGACATCCAGACG	AGTICAAACACAACTGAT
91001192	pml1653	2,8,12	TGCAATAMGGAAAGCCA	CGGTGTTAGGTTAGATG
91000213	pml1776	2,9,13,17,X	GTCATTGTGTCATTTCC	ACATTTATTTTCAACG
91000919	pml0985	20,X	CATGACACAGGGCACITC	GCAACTACAAATGCCAACT
91001169	pml0157	3,10,15	CAGGAACTGGAGGAAAG	GATTAACTGAGAAAG
91000071	pml2651	3,4,M	TTAGGAAATAGGTTAGAG	ATAGTATGGTTGACACGTA
91001128	pml2632	3,6	TGGATTTGCTTACCTGTT	ACACCCTGAGGTTTAC
91001191	pml1123	4,8	GCACACAGGCAACACAGA	CCTCTAACACACACGAG
91000217	pml2250	3,9,10,15	GGATTCTTATGGCTCAT	GTTATGTTGACGCAATTAC
91000605	pml0626	4,6	GCATTAAACGAAACATA	CTTCACGAGGCAAAAC
91001212	pml1214	6,20	AGAIGCTAACATTAGGATA	TTTATGACATAGAGGAGT
91001112	pml0606	7,18	CCAGACTACGGCTGATGGC	CCCTTACCCAGAACACIY
91001111	pml1253	9,11	ACCAGTCACGCTCTAAATA	CCCATATAAGTGGAGGATGTC
91002357	pml0115	9,11	AAGAATGTTTACGTT	TTATGACTGGGAAAT
91001261	pml1224	10,15,22	ACTACCTGAGATATTAGT	TCTATTGTTGATGTTGTA
91001154	pml2120	11,M	ATACCACTTCGCTTCAGG	GAGGGAGCTTCAGGCTCT
91002320	pml2203	12,19	GCACCAAGAGACATTCAG	TGGGAAATGAAATAACT
91002114	pml2613	12,M	GATCTCAGTCCTGGTTATT	TACATACMAGGAAACAGT
91001103	pml2273	12,M	AATCTTGTTGCTGCTTC	GTCCTCTCTGAGGCTGA
91001487	pml3225	13,16	AACCTGTTTACCGCATCTT	AGGTTATGTCACCAAGAA
91002316	pml2780	14,16	TGTGTTACCATGAGAC	AGMACACACATCAAATGC
91001435	pml1683	17,20,C	GAATGTCATCCAGAGTAG	CTAGTATATCCGGCTCG
91001391	pml1746	17,22,Y	TITATECCAGCAAGCAAC	TCTCTCTCTACTTCCTTC
91000556	pml0964	17,C	ACTTAAGTAGCTTGTACG	TGCTCTCTGCTGCTGATA
91001389	pml2217	17,C	CCCAGTTAAAGATATTGT	ATGAGGATGGAGGAGTGA
91001140	pml213	18,C	TGCAAGAGGATTTCCAG	CTAGGGTCATCTTCAGC
91001217	pml1116	19,20	ATCCCTGCTTACTOACTCAG	GCCTGTTAACOACTCAG
91000099	pml3024	19,22	GCCTACATCTGTTGACT	ACCTCTGGAAACAACTAT
91001172	pml0887	19,22		

Fig. 10

91001057	pm12049	C	AGGACACAAACACCAAGCTAT	TTCCTGATTAGACATGAC	
91001177	pm1153	M	ATCTCTTGTAGCCATCTA	GTTAACGGCTGTGCCATT	
91001056	pm12216	M	GTAGAGCTGCATGACIACC	ACAGACAAAGATAATCATA	
91001168	pm10506	N	GTCCCCACAGGCCACCTAAC	CCACACAAATTAGATCCATC	
91001151	pm12354	N	TGTCCTTGGAACCTGCCT	TITAAACGTCATATAATAGTT	
91000229	pm12492	M,C	GTCTGAAGAGGGGACTICA	CITAACTCGATAGCCAGTC	
91000231	pm12746	M,C	CAAAACACGCAAACCTAC	ATGGTTTAATTACAGATQ	
91000235	pm11704	M,C	TCCACCCAGAAAGCACACT	ATTCATAGGGAAAGGTC	
91000302	pm12316	M,C	TCGAGAAGGACAAATCACC	GAACAGGGTTAGTCATTCG	
91000541	pm1689	M,C	CATGGGCTACGGAAACAGG	AGGACCTCGGGGCTCTGAG	
91000615	pm11442	M,C	AAAGGATCTGAGAGGAAACA	GGAGGACCTGGCTGGCTCTA	
91000722	pm1452	I,A,C	GGAGCAQATACCTTACACC	TGGTCATTAGTCCTTC	
91000995	pm1268	M,C	GAAGGCTCTGGAGGAAGT	CAGACCCCATTTTATAGC	
91001016	pm12703	M,C	ACGATATTATAGCTAGTA	TCAAACACTTAAATATGCT	
91001031	pm1144	M,C	AGATAGCTGGGGCTCAGAA	CCATCCCGTCAATCCAGTT	
91001127	pm12290	M,C	ACTGTCGATGGAAAGTTACA	CCACACGIGGACCCGCT	
91001167	pm1626	M,C	GADACCCCTGTCATCTTA	CTTCCTCTGGCTCTCTGT	
91001216	pm12109	M,C	TAGTCAGATTCAGTAGT	ACATGTTTTTGTAGTCT	
91001231	pm11210	M,C	AACGTTCTCCATCAAGCTG	ATGTTAAACCTCTCACC	
91001281	pm1121	M,C	ACTAAAAACCCACCGCAT	ACACAGCAGTCATAAGAA	
91001315	pm0952	M,C	AGAAGGAGTTCTGGCTCTA	ATCTGGAGAGGGGAGGA	
91001356	pm12116	M,C	ATCTGATGACCTATATCT	CGTCCTCTTATTTGACAT	
91001411	pm0958	M,C	ATGGGTTTATGGGGGTTTC	GAGACCAAGGACCTCTA	
91001420	pm12626	M,C	ACATYNAAGGGAGTAAGGT	GGACATTCATGCCACAGC	
91001432	pm1210	M,C	TGTTGAGCTGGCTTCTGAA	CAGGCCCTCTGACTGAGACA	
91001552	pm10109	M,C	GCCCCAGAGACATCTCT	TCTTAAAGGGCTCTGGTG	
91001664	pm12012	No product	CAACCACTTGGGGTAAGT	GAATTAACCTCTGCTCTA	
91000050	pm10204	No product	CTTGGGATATTCCTCAT	CCCTGGGGTACTTCTAAG	
91002611	pm0968	No product	AGCCAGCCCTCTGTAIG	CTGGATTGTTCTGATCTG	
91001152	pm11622	No product	TGTTGAGTGAAGATACTG	TTTGTATGAGACACACT	
91001154	pm11615	No product	CAGTAGCTGGCTTGAATG	TTATGIGAAATGGCTGT	
91001165	pm2908	No product	TACACCCACTCTAAAGTC	TTTGGGATCAAGGAATCT	
91001171	pm10261	No product	TACATTCTCAGACCTCTG	TTTCAAACACTTATCTT	
91001556	pm0849	No product	ATCAAGCTCAGTCATCG	ATTTCGCCCTCTGATGGIC	
91001571	pm1281	No product	GAICTTGGCCCTTAATGGA	TTTGGAGTCACTTATTC	
91001622	pm1606	No product	GATCTCCTGTCCTTACA	TTTAAACAGACACATAC	
91001640	pm0832	No product			
			26	45	45

Fig. 11

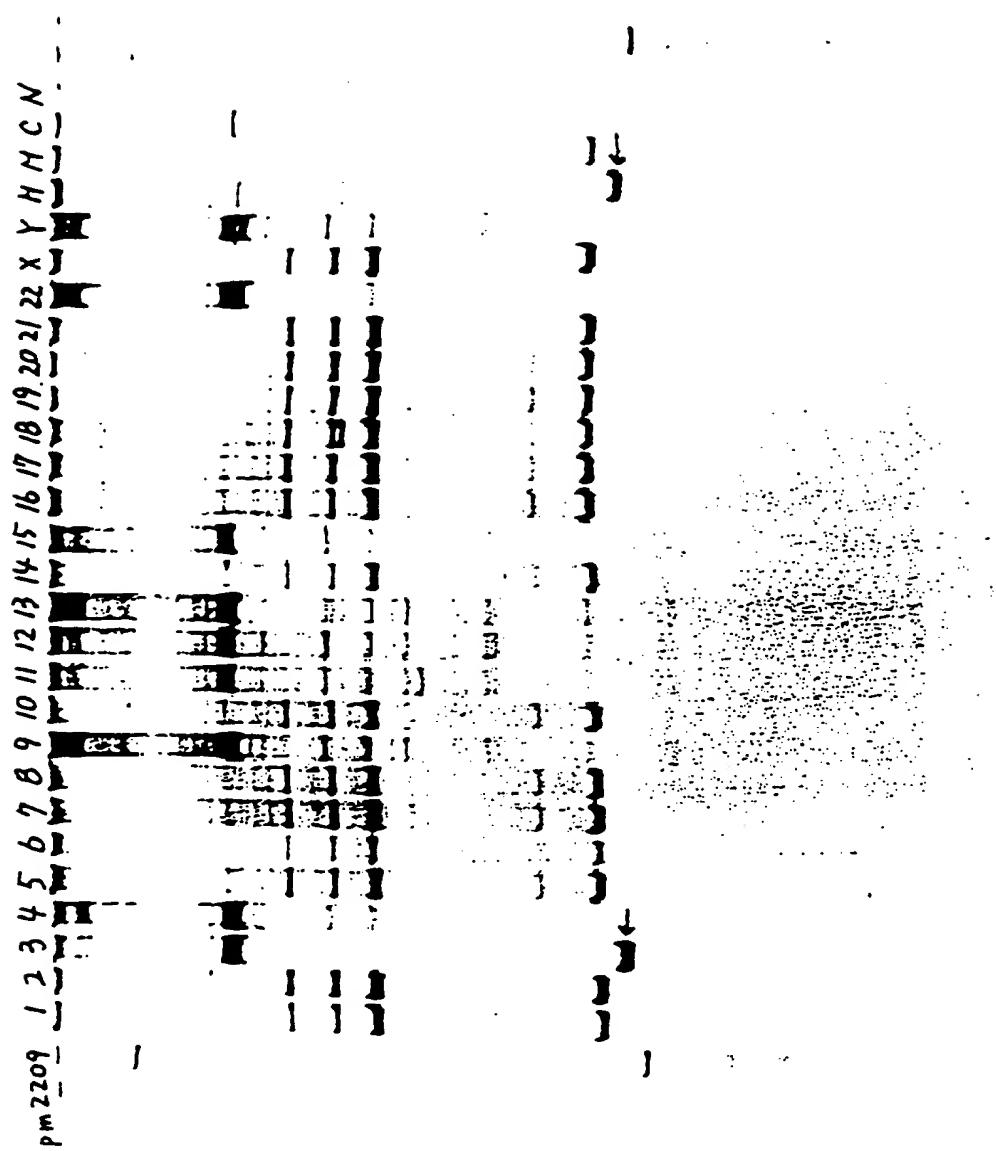


Fig. 12

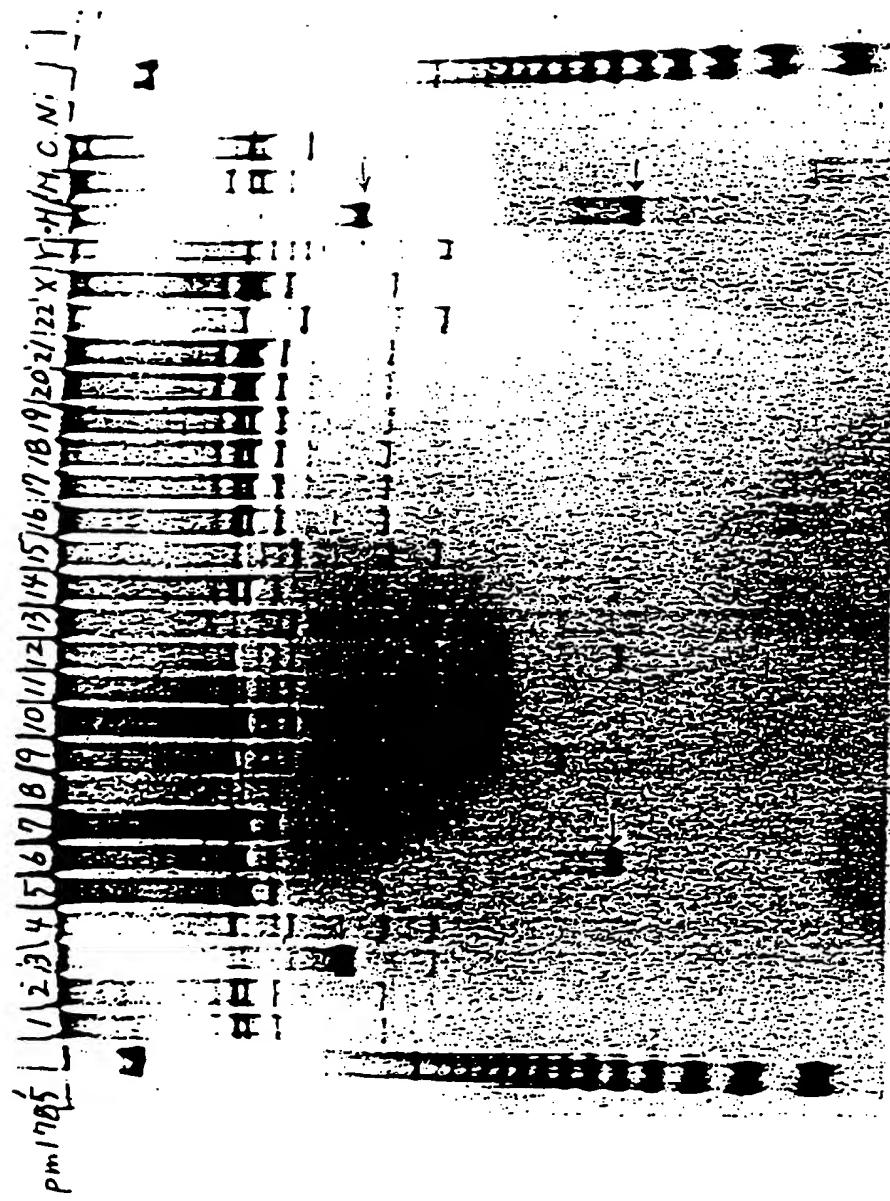


Fig. 13

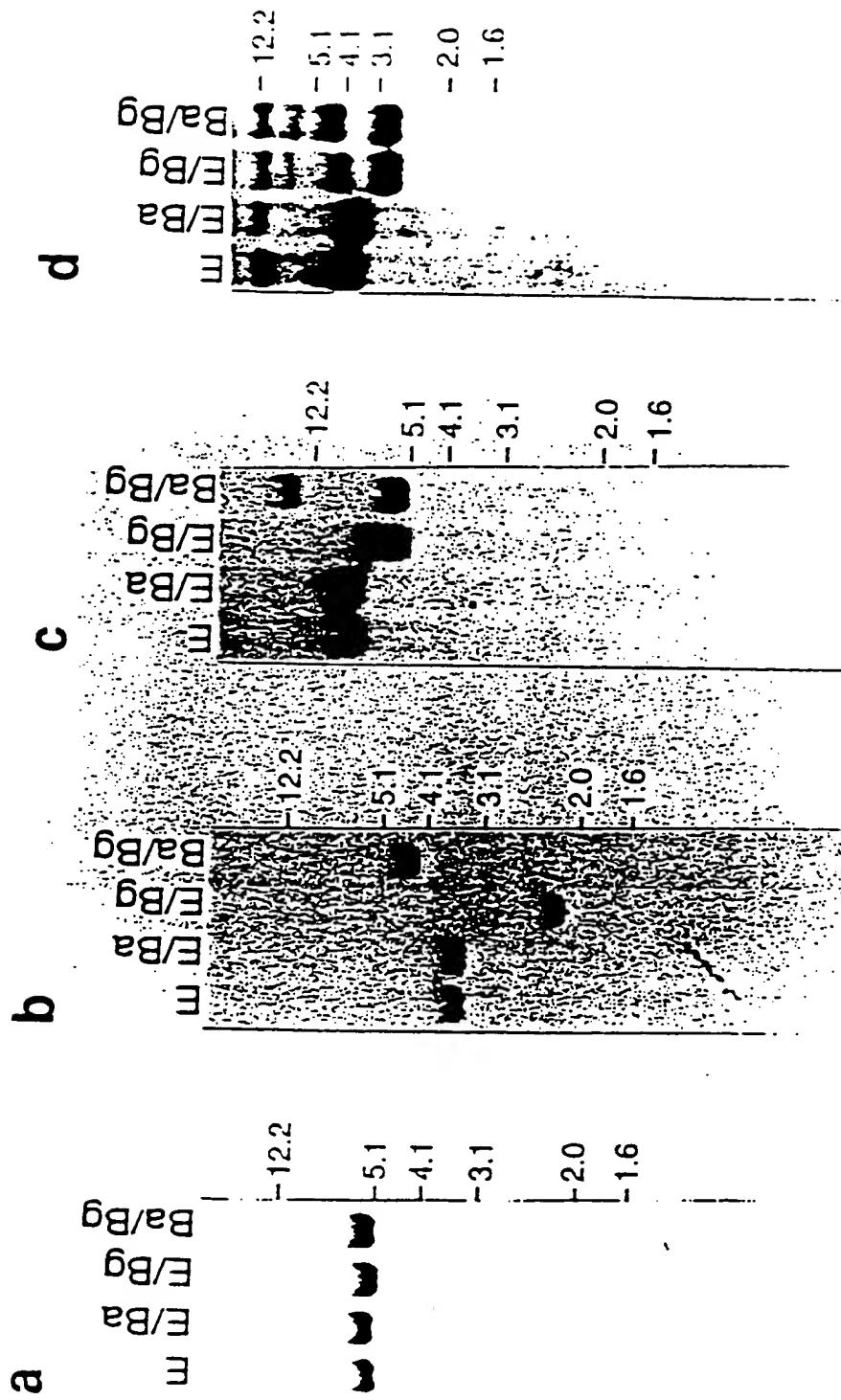


Fig. 14

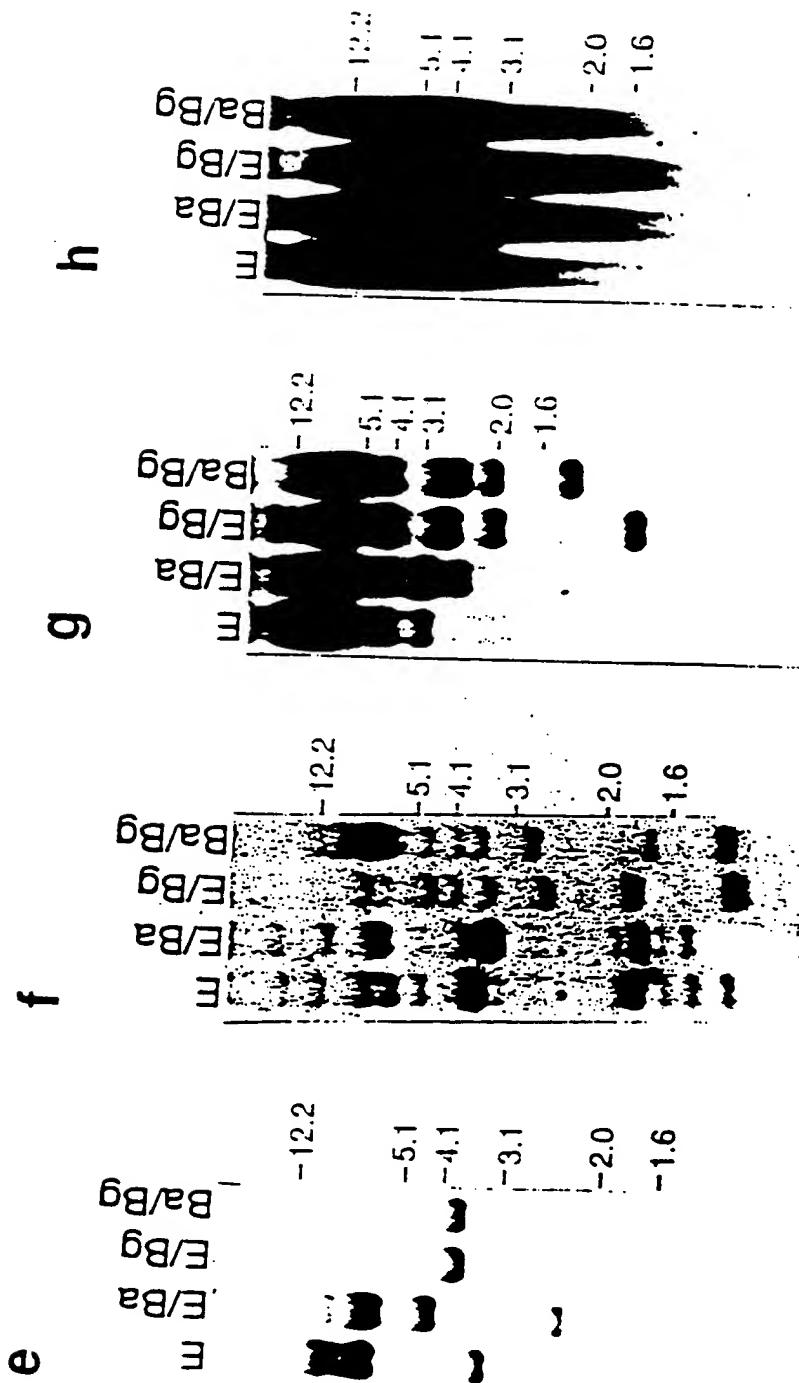


Fig. 15

Hybrid cells used for Southern hybridization

Hybrid cell	Human chromosome No.	Parent cell	Intact chromosome (%)	Translocated chromosome (%)
A9(neo-1)-4	1	A9	100 (0)	0
A9(neo-2)-1	2	A9	93 (3)	0
GM10253	3	CHO	100 (0)	0
GM10115	4	CHO	100 (0)	0
A9(neo-5)-4	5	A9	40 (0)	90
A9(neo-6)-3	6	A9	100 (60)	0
A9(neo-7)-2	7	A9	100 (89)	0
A9(neo-8)-1	8	A9	91 (82)	0
GM10611	9	CHO	79 (5)	11
A9(neo-10)-3	10	A9	94 (6)	75
A9(neo-11)-1	11	A9	24 (0)	76
GM10927A	11	CHO	96 (21)	4
A9(neo-12)-4	12	A9	0 (0)	100
GM10868	12	CHO	82 (6)	0
GM10898	13	CHO	82 (0)	10
GM10479	14	3T6	76 (29)	0
A9(neo-15)-2	15	A9	9 (0)	78
GM11418	15	CHO	62 (0)	100
GM10567	16	A9	69 (0)	0
GM10496	17	LTMK	80 (10)	0
A9(neo-18)-5	18	A9	100 (66)	0
A9(neo-19)-1	19	A9	92 (23)	8
A9(neo-20)-3	20	A9	61 (5)	17
GM08854	21	A9	81 (24)	0
GM10027	22	CHO	93 (0)	100
GM10324	X	A9	61 (10)	0
GM06317	Y	CHW1103	91 (0)	9

Fig. 16

a
Chromosome

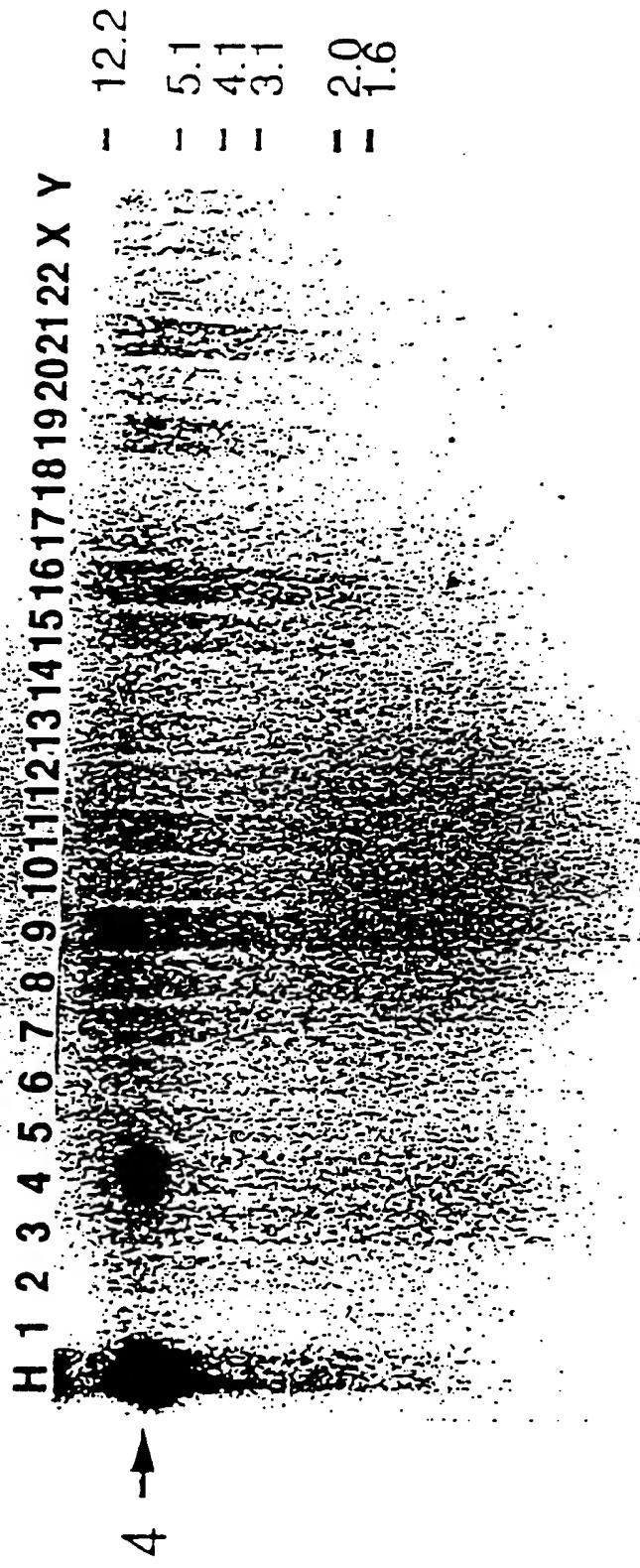


Fig. 17

b

Chromosome

H 1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 X Y

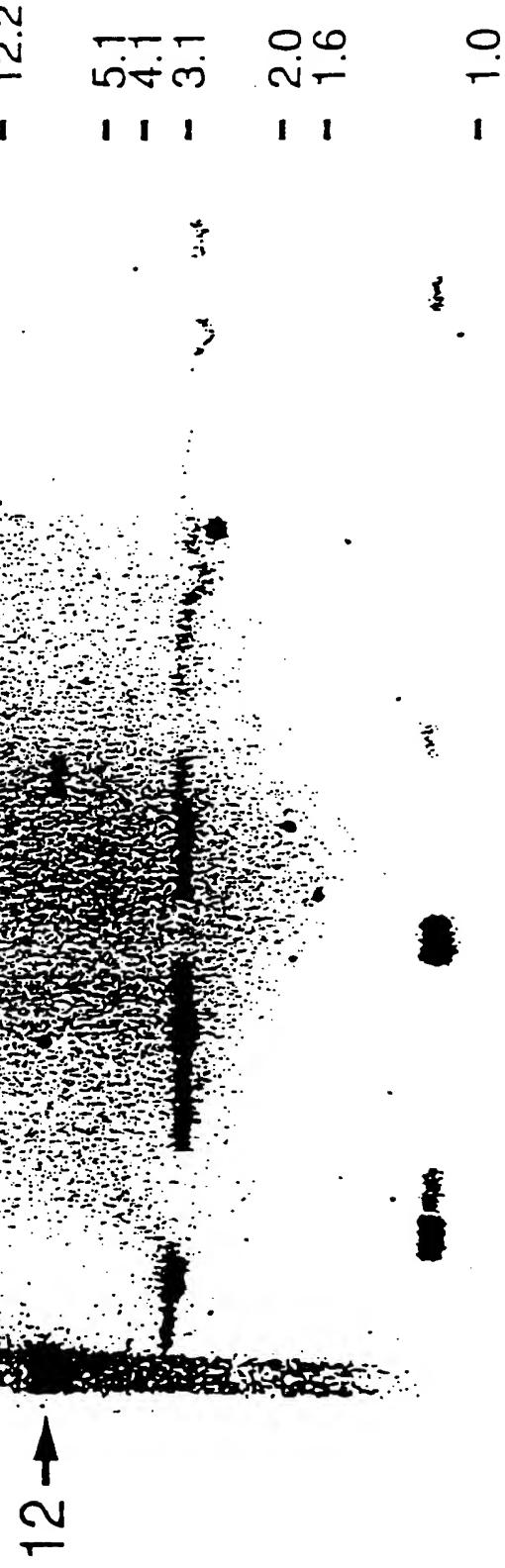


Fig. 18

C

Chromosome

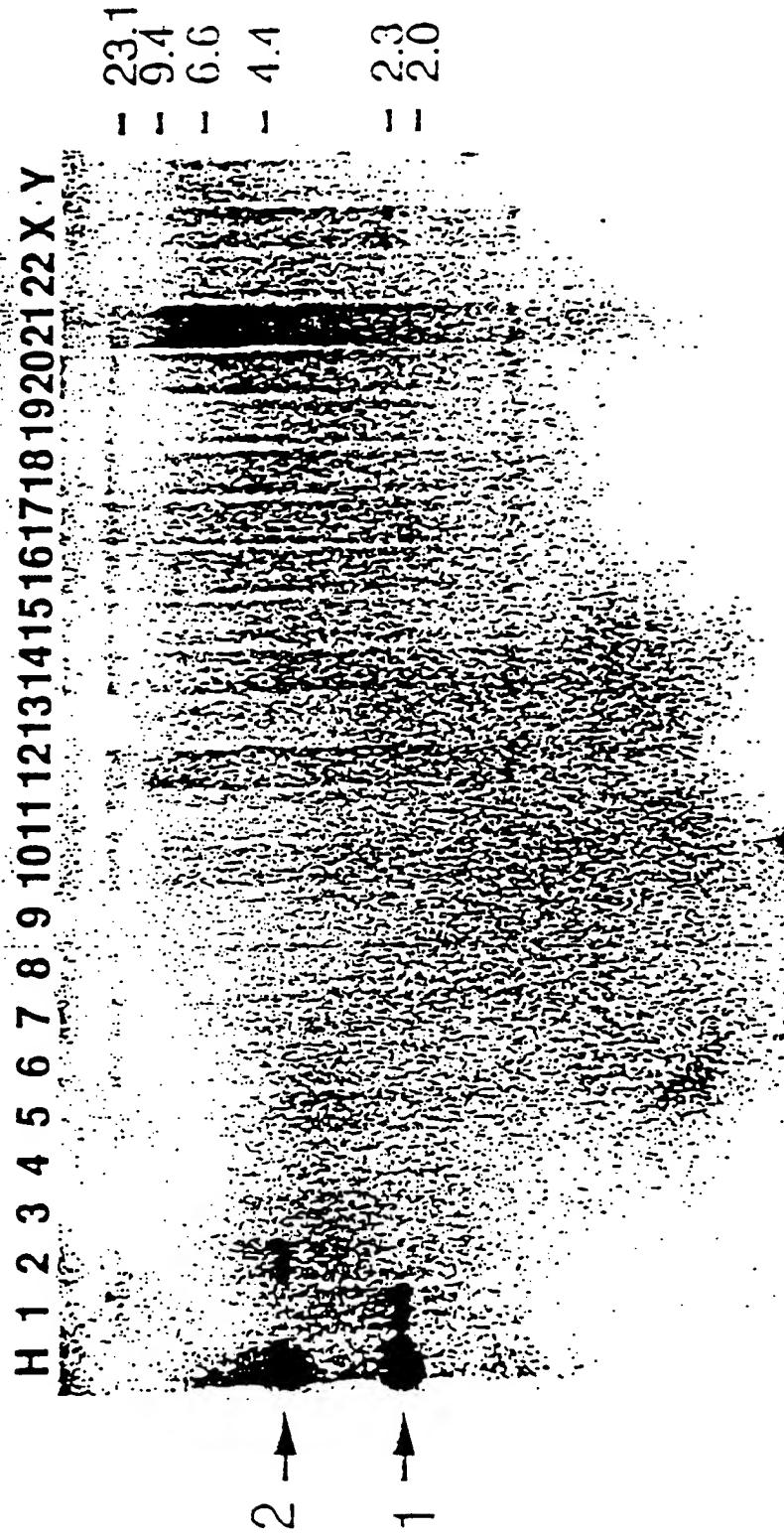
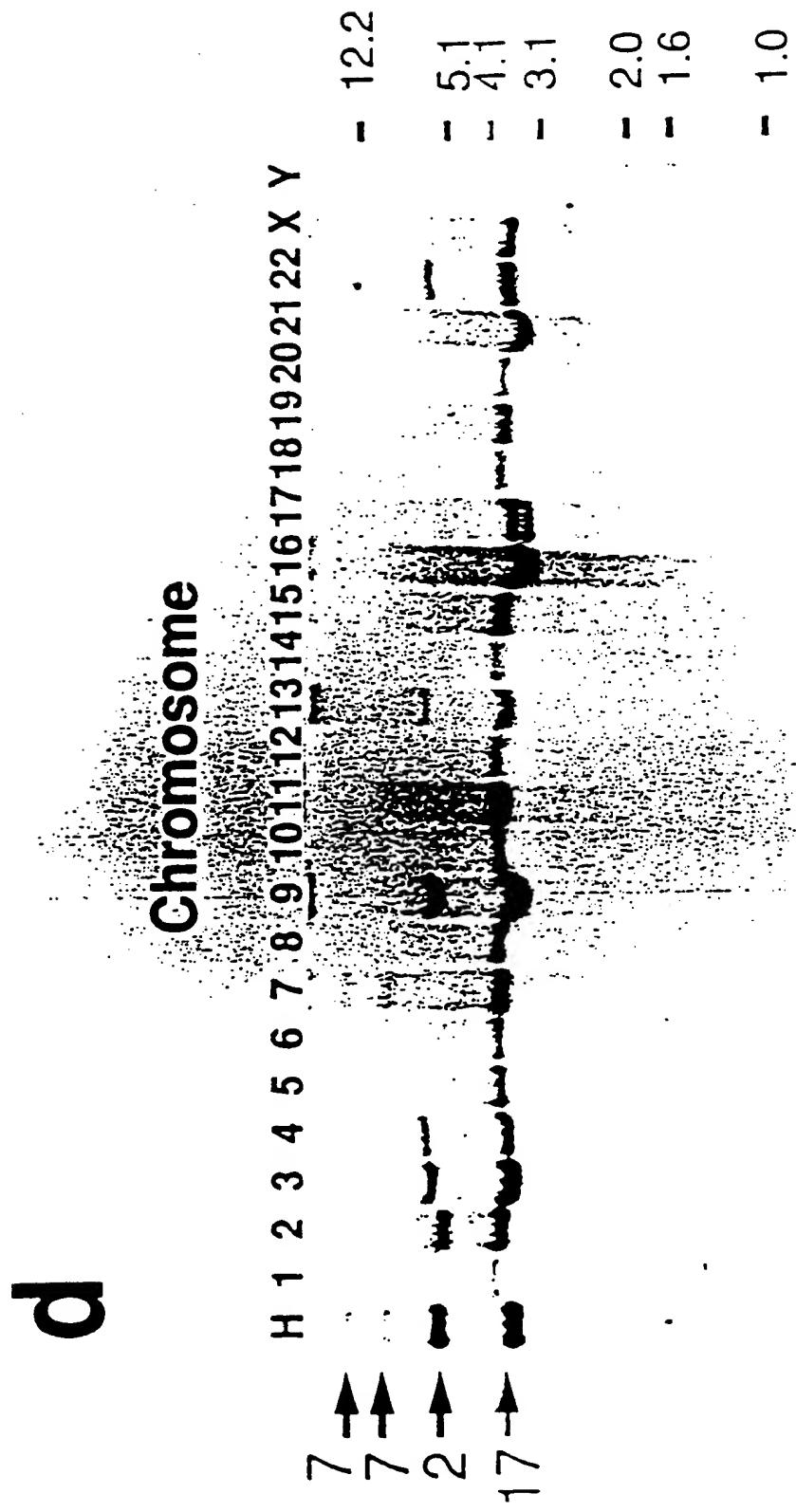


Fig. 19



e

Chromosome

H 1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 X Y

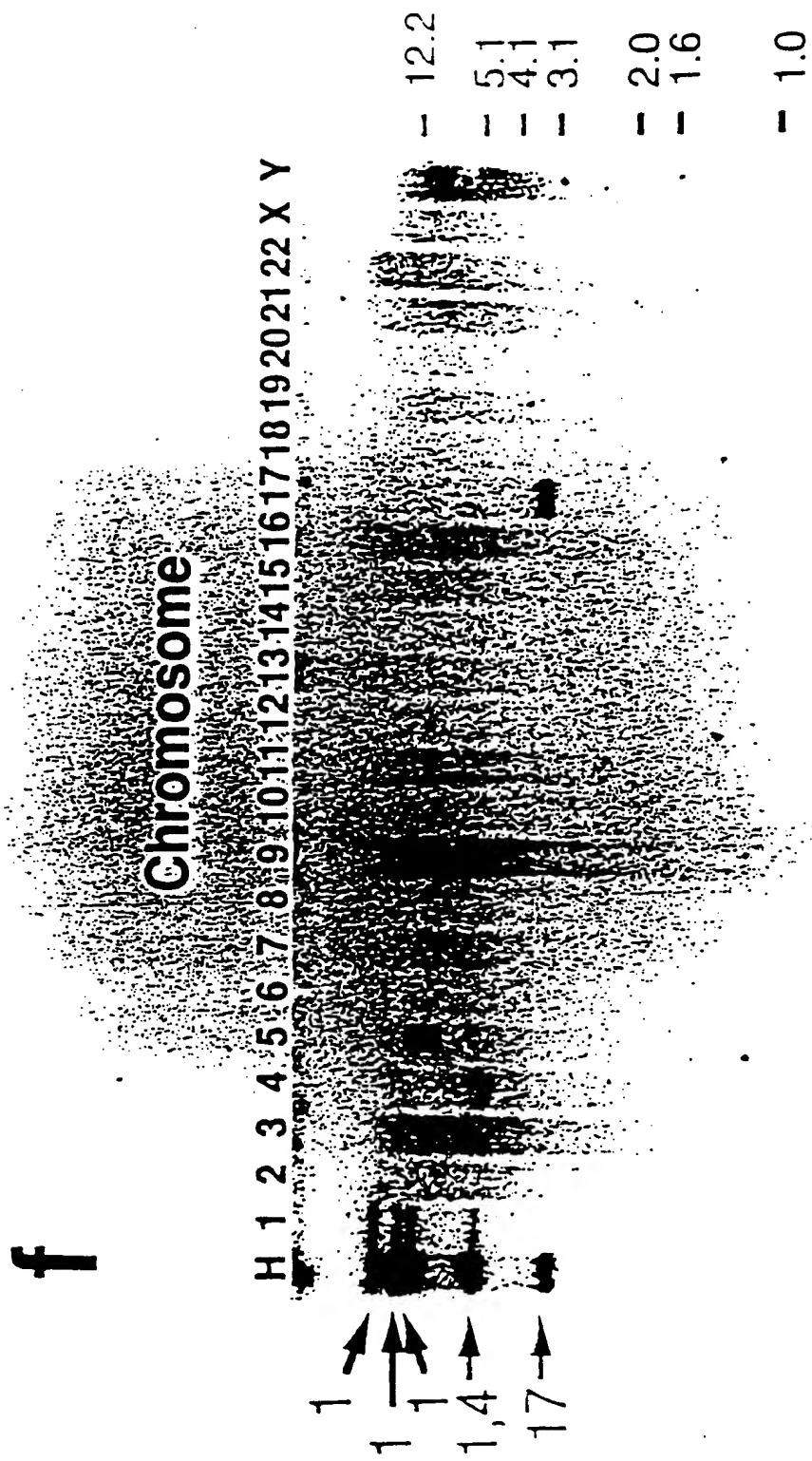
7,19
8
11,12
11



Fig. 20

EP 0 679 716 A1

Fig. 21



G

Chromosome

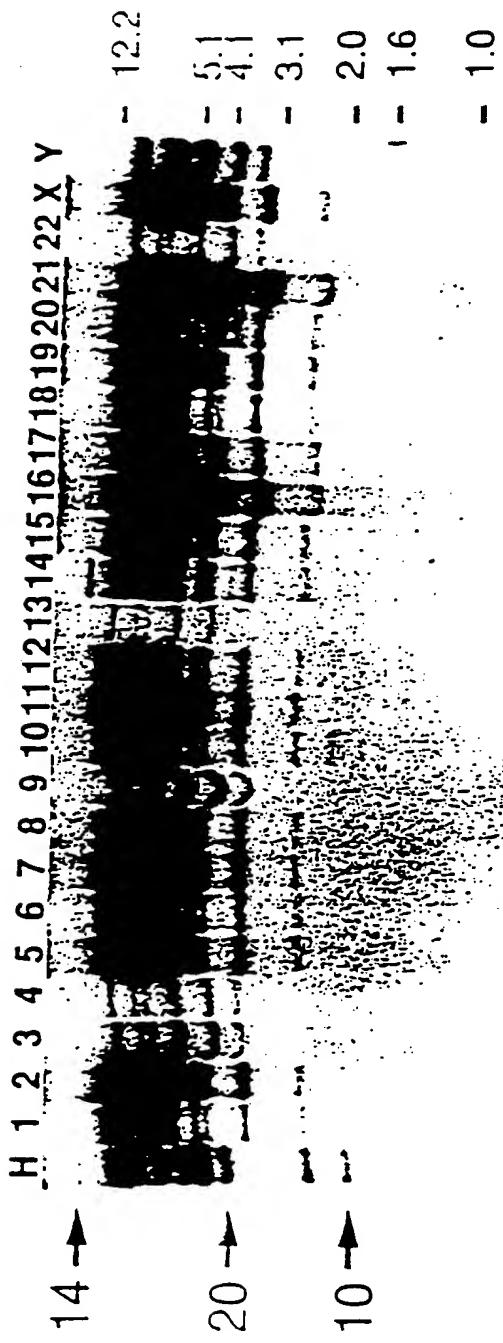


Fig. 22

Fig. 23

Chromosomal mapping of each GS by Southern blot technique

Clone	Sequence length	Numbers of bands detected with human whole chromosomes			Chromosomes assigned	Background	
		2/3	2/3; 3/3	Mouse		Chinese hamster	
Single band group:							
c12e11	GS000073	432	1	1	1	9	0 0
c12e06	GS000062	540	1	1	1	6.15	0 0
c12g01	GS000280	212	1	1	1	2	1 1
c13c05	GS000117	359	1	1	1	11-	0 0
c13c07	GS000129	355	1	1	1	2	0 0
c13f10	GS000206	267	1	1	1	14	0 0
c13h01	GS000273	133	1	1	1	12-	0 0
c13h02	GS000222	167	1	1	1	6	0 0
d0gG2	GS000095	397	1	1	1	3	0 0
d0h07	GS000164	313	1	1	1	11	1 1
d1510	GS000348	153	1	1	1	20	0 0
hm01e12	GS000223	246	1	1	1	27	0 0
hm01e09	GS000423	157	0	1	1	1	0 0
hm01e12	junk	394	1	1	1	17	0 0
hm01e05	GS000068	454	1	1	1	19.22	0 0
hm01e10	GS000299	173	0	1	1	10	0 0
hm01g09	GS000053	477	1	1	1	6	0 0
hm01h07	GS000115	363	1	1	1	12	0 0
hm02a02	GS000130	344	1	1	1	4	0 0
hm02a04	GS000329	164	1	1	0	10	0 0
hm02e01	GS000293	271	1	1	1	16	0 0
hm02e01	GS000016	590	1	1	1	20	0 0
hm02e02	GS000342	156	0	1	1	14	0 0
hm02e05	GS000401	223	1	1	0	n.d.	0 0
hm02g02	GS000191	273	1	1	1	17	0 0
hm05e05	GS000251	219	1	1	1	6	2 0
hm05a10	junk	392	1	1	1	1	1 1
hm05c10	GS000009	606	1	1	1	1	0 0
km001	junk	169	1	1	0	n.d.	0 0
s10S	GS000001	703	1	1	1	5	0 0
s110	GS000057	471	1	1	1	3	0 0
s11d11	GS000307	#175	0	0	1	7	0 0
s11h01	GS000299	204	1	1	1	3	0 0
s147	GS000080	461	1	1	0	2	0 0
s14e06	junk	639	1	1	1	1	0 0
s14g02	GS000152	322	1	1	1	2	0 0
s14h12	GS000271	193	1	1	1	4	1 1
s150	GS000143	330	1	1	1	17	0 0
s15G	GS000002	306	1	1	1	2	1 1
s15b11	GS000250	221	1	1	1	14	0 0
s179	GS000275	196	1	1	1	n.d.	0 0
s246	GS000234	241	1	1	1	9	0 0
s247	GS000247	153	1	1	1	1	0 0
s270	junk	135	1	1	1	19	0 0

Fig. 24

Clone	Numbers of bands detected with human whole chromosomes						Chromosomes assigned	Background	
	Sequence length	Σ	$\Sigma/3_1$	$\Sigma/3_2$	$\Sigma/3_3$	$\Sigma/3_5$		Chinese Mouse	hamster
s306	GS000256	205	1	1	0	1	X	0	0
s309	GS000171	305	1	1	0	1	1	0	0
s342	GS000323	165	1	1	1	1	4	3	2
s331	GS000265	207	1	1	0	1	6.15	1	1
s334	GS000165	112	1	1	1	1	1	0	0
s337	GS000276	195	1	1	1	1	17	0	0
s339	GS000295	130	1	1	1	1	n.d.	0	1
s443	GS000330	251	1	1	1	1	n.d.	0	0
s470	junk	251	1	1	1	1	17	0	0
s474	GS000192	273	1	1	1	1	5	0	0
s503	junk	312	1	1	1	1	12	0	0
s507	junk	600	1	1	1	1	1	2	1
s517	GS000334	161	1	1	1	1	14	1	1
s632	junk	337	1	1	1	1	2	0	0
s633	GS000186	311	1	1	1	1	22	2	1
s550	GS000041	644	1	1	1	1	12	1	1
tw1-04	GS000025	537	1	1	1	1	3.7	0	0
tw1-19	GS000218	255	1	1	1	1	17	0	0
tw1-32	junk	250	1	1	1	1	5	0	0
tw1-37	GS000237	235	1	1	1	1	22	0	0
tw1-42	junk	391	1	1	1	1	8	1	1
tw1-43	GS000096	178	1	1	1	1	14	0	0
tw1-96	GS000133	339	1	1	1	1	11	0	0
Two band group :									
c12f12	GS000195	277	1	2	2	2	1,	1	1
c13d02	GS000042	503	2	2	1	1	2,	0	0
hm01a06	GS000129	344	2	2	2	2	11.13	3	5
hm01a07	GS000207	269	2	2	2	2	7,	0	0
hm01d05	GS000232	243	2	2	2	1	2,	0	0
hm01e01	GS000181	292	2	2	2	2	1.2	0	0
hm02a08	GS000435	302	2	2	2	2	3,	1	1
hm02c04	GS000221	253	2	2	2	2	3,	0	0
hm02c05	GS000146	332	2	2	2	2	17.19.22	0	0
hm03f07	GS000043	503	1	1	2	1	3,	0	0
s11d06	GS000268	205	2	2	2	2	11.12	0	0
s11g12	GS000337	255	2	2	2	2	6,	0	0
s124	GS000083	404	2	2	2	2	9,	1	1
s144	GS000132	342	1	2	2	2	1.7	0	0
s14f03	GS000239	243	1	2	2	2	2,	3	2
s15e02	junk	439	2	2	1	2	6,	0	0
s16b09	junk	420	1	1	1	2	10.14	0	0
s17c09	GS000248	223	2	2	2	2	14,	0	0
s231	junk	234	2	2	2	2	11,	0	0
s254	GS000124	151	2	2	2	2	1,	3	1
s255	GS000235	219	2	2	2	2	11,	0	0
s272	junk	195	2	2	2	2	10.16	1	1

Fig. 25

Clone	Sequence length	Numbers of bands detected with human whole chromosomes					Chromosomes assigned	Background	
		E	E/B ₁	E/B ₂	E/B ₃	Background		Mouse	Chinese hamster
s311	GSJ00092	333	1	1	2	2	16.	1	1
s313	junk	131	2	2	1	0	20.	0	0
s317	GSJ00100	339	0	0	1	2	14,14	1	1
s336	GSJ00134	337	2	2	2	2	12,14	0	0
s333	GSJ00139	233	2	2	2	1	22,X	0	0
s339	GSJ00233	137	2	1	1	2	17.	0	0
s394	GSJ00063	449	2	1	2	2	13,14	0	0
s396	junk	277	2	2	2	2	17.	0	1
s455	junk	452	1	2	2	1	4.	0	0
s456	GSJ00236	132	2	2	2	2	8,10	1	2
s465	GSJ00201	274	1	1	2	2	6,15	0	0
s635	junk	250	1	1	1	2	9,13	0	0
s639	GSJ00257	205	1	2	2	2	2X	0	0
s656	GSJ00025	#590	2	2	0	2	6,11	0	0
tw1-33	junk	352	2	2	2	2	1.	0	0
tw1-39	GSJ00153	#321	2	2	2	2	17.	0	0
tw1-70	GSJ00061	441	1	1	2	1	11.	0	0
tw1-80	junk	453	2	2	1	2	9,17	2	2
tw1-87	GSJ000158	316	2	2	2	2	7.	0	0
Three band group									
d0h06	GSJ00020	417	3	3	3	1	1.	0	0
hm05b07	junk	336	2	3	3	3	5.	0	0.
hm05g02	GSJ000209	267	2	2	2	1	3,17,19	1	1
s129	GSJ000107	373	3	3	3	3	n.d.	1	1
s173	GSJ000357	146	1	2	2	3	2.	0	0
s17a10	GSJ000294	131	3	3	3	3	2,13,22	1	1
s302	GSJ000412	638	2	2	2	3	XCC	1	1
s401	GSJ000224	249	2	1	3	3	6,6.	0	0
s654	GSJ000045	491	3	3	3	3	1,22.	0	0
tw1-82	GSJ000208	267	3	3	3	3	13.	4	0
Four band group									
c12g07	GSJ00154	320	4	4	2	3	5, 14,	0	0
c13a08	GSJ00055	508	3	3	4	4	2,7,7,17	1	2
c13c04	GSJ000106	#376	4	3	3	3	n.d.	0	2
c13e09	GSJ000302	195	4	2	4	4	2,17,	7	2
s136	GSJ000160	315	4	4	4	4	4X.	2	1
s163	GSJ000004	#618	4	4	4	2	4,4,8,20	3	1
s479	GSJ000130	293	4	4	2	2	7,8,11,11,12,19	0	0
Group with 5 or more bands									
c12f08	GSJ000253	217	5	5	5	2	2,7,9,14,	2	0
hc01	junk	374	12	12	15	13	1,2,6,	22	20
hd10	junk	361	4	4	4	8	n.d.	12	6
he10	junk	173	6	2	3	3	6,3,9,19,21,	3	3
hm01c05	GSJ000205	176	9	7	5	5	X	9	8
hm01f04	GSJ000246	215	8	10	5	5	n.d.	12	12
hm01g02	junk	411	9	6	6	4	10,14,20,	14	6

Fig. 26

Clone	Sequence length	Numbers of bands detected with human whole chromosomes					Chromosomes assigned	Background	
		E	E/B ₁	E/B ₂	B ₁ /B ₂	Mouse		Chinese hamster	
hm02f09	GS000273	442	9	7	7	5	3,3,6,11,13,14,15,16	0	0
hm05z02	GS000096	373	5	6	4	6	2,3,17,	3	3
hm05z04	GS000236	#239	6	6	6	7	n.d.	3	5
kmb01	junk	350	3	5	5	5	13,	14	7
s11f06	GS000319	170	6	6	6	4	1,2,2,3,4,6,13,15,	0	3
s14f01	GS000407	262	12	11	10	9	1,5,9,13,	6	3
s173	GS000094	397	5	4	6	3	1,1,1,1,4,17	0	0
s255	GS000323	167	10	12	11	14	13,	9	5
s341	junk	494	9	9	8	6	n.d.	15	3
s406	GS000113	364	6	7	5	4	2,7,8,13,20,20	4	1
tw1-46	junk	593	9	10	10	10	1,1,2,5,11,X,	3	5
tw1-63	junk	203	3	10	10	12	3,4,	17	11
Bands no detected:									
c13g02	GS000340	157	0	0	0	0	-	-	-
hm0le10	junk	232	0	0	0	0	-	-	-
hm02d11	GS000274	196	0	0	0	0	-	-	-
s323	GS000273	194	0	0	0	0	-	-	-
s359	GS000199	279	0	0	0	0	-	-	-
s511	junk	233	0	0	0	0	-	-	-
s645	GS000012	#734	0	0	0	0	-	-	-
s647	GS000105	360	0	0	0	0	-	-	-
s651	junk	510	0	0	0	0	-	-	-

INTERNATIONAL SEARCH REPORT

International application No.

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A. CLASSIFICATION OF SUBJECT MATTER

Int. Cl⁶ C12N15/11, C12Q1/68//G01N33/566

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

Int. Cl⁶ C12N15/11, C12Q1/68//G01N33/566

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

BIOSIS PREVIEWS, CAS ONLINE

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	Nucleic Acids. Res., Vol. 15, 1987, Ou, J. H. "Cloning and characterization of a human ribosomal protein gene with enhanced expression in fetal and neoplastic cells" p. 8919-8934	1-6 (164)
X	Differentiations, Vol. 33, 1986, Oshima, R. G. et al. "Comparison of mouse and human keratin 18:A component of intermediate filaments expressed prior to implantation" p. 61-68	1-6 (226)
X	J. Biol. Chem., Vol. 265, 1990, Wilkin, D. J. et al. "Isolation and sequence of the human farnesyl pyrophosphate synthetase cDNA: coordinate regulation of the mRNAs for farnesyl pyrophosphate synthetase, 3-hydroxy-3- methylglutaryl coenzyme A reductase, and 3- hydroxy-3-methylglutaryl coenzyme A synthetase" p. 4607-4614	1-6 (255)

 Further documents are listed in the continuation of Box C. See patent family annex.

* Special categories of cited documents:

- "A" document defining the general state of the art which is not considered to be of particular relevance
- "E" earlier document but published on or after the international filing date
- "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- "O" document referring to an oral disclosure, use, exhibition or other means
- "P" document published prior to the international filing date but later than the priority date claimed

"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention

"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone

"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art

"&" document member of the same patent family

Date of the actual completion of the international search
February 6, 1995 (06. 02. 95)Date of mailing of the international search report
March 7, 1995 (07. 03. 95)Name and mailing address of the ISA/
Japanese Patent Office
Facsimile No.

Authorized officer

Telephone No.

INTERNATIONAL SEARCH REPORT

International application No.

PCT/JP94/01916

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Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
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X	J. Biol. Chem., Vol. 263, 1988, Fischer, R. et al. "Multiple divergent mRNAs code for a single human calmodulin" p. 17055-17062	1-6 (386)
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X	J. Biol. Chem., Vol. 265, 1990, Natsumeda, Y. et al. "Two distinct cDNAs for human IMP dehydrogenase" p. 5292-5295	1-6 (454)
X	Genes Dev., Vol. 7, 1993, Patton, J. G. et al. "Cloning and characterization of PSF a novel pre-mRNA splicing factor" p. 393-406	1-6 (706)
X	Nucleic Acids Res., Vol. 16, 1988, Stanford, D. R. et al. "The complete primary structure of the human snRNP E protein" p. 10593-10605	1-6 (711)
X	Proc. Natl. Acad. Sci. U.S.A., Vol. 84, 1987, Inoue, C. et al. "Evolutionary conservation of the insulinoma gene rig and its possible function" p. 6659-6662	1-6 (723)
X	J. Immunol., Vol. 144, 1990, Jongstra-Bilen, J. et al. "Human and mouse LSP1 genes code for highly conserved phosphoproteins" p. 1104-1110	1-6 (741)
X	Biochem. J., Vol. 248, 1987, Sakai, I. et al. "The cDNA and protein sequences of human lactate dehydrogenase-B" p. 933-936	1-6 (772)

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Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
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X	Mol. Cell. Biol., Vol. 3, 1983, Cowan, N. J. et al. "Expression of human alpha-tubulin genes: interspecies conservation of 3' untranslated regions" p. 1738-1745	1-6 (820)
X	Nucleic Acids Res., Vol. 17, 1989, Taaman, J. W. et al. "Nucleotide sequence of cDNA encoding subunit VIb of human cytochrome c oxidase" p. 1766-1766	1-6 (844)
X	Gene, Vol. 93, 1990 Taanman, J. W., Schrage, C., Ponne, N., Das, A., Bolhuis, P. A., de Vries, H. and Agsteribbe, E. Isolation of cDNAs encoding Subunit VIb of human cytochrome c oxidase and steady-state levels of coxVIb mRNA in different tissues p. 285-291	1-6 (844)
X	J. Biol. Chem., Vol. 264, 1989, Gray, P. W. et al. "Cloning of the cDNA of a human neutrophil bactericidal protein: Structural and functional correlations" p. 9505-9509	1-6 (861)
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X	Science, Vol. 248, 1990, Smith, C. A. et al. "A receptor for human tumor necrosis factor defines an unusual family of cellular and viral proteins" p. 1019-1023	1-6 (1431)

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X	J. Clin. Invest., Vol. 83, 1989, Look, A. T. et al. "The human myeloid plasma membrane glycoprotein CD13 (gp150) is identical to aminopeptidase N" p. 1299-1307	1-6 (1469)
X	J. Cell Biol., Vol. 105, 1987, Argraves, W. S. et al. "Amino Acid Sequence of the Human Fibronectin Receptor" p. 1183-1190	1-6 (1607)
X	Nucleic Acids Res., Vol. 18, 1990, Liebhaber, S. A. et al. "Characterization of a human cDNA encoding a widely expressed and highly conserved cysteine-rich protein with an unusual zinc-finger motif" p. 3871-3879	1-6 (1642)
X	J. Biol. Chem., Vol. 264, 1989, Didsbury, J. et al. "Rac, a novel ras-related family of proteins that are botulinum toxin substrates" p. 16378-16382	1-6 (1709)
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X	J. Biol. Chem., Vol. 266, 1991, Wu, Y. et al. "Activation of globin gene expression by cDNAs from induced K562 cells: Evidence for involvement of ferritin in globin gene expression" p. 17566-17572	1-6 (1785)
X	Proc. Natl. Acad. Sci. U.S.A., Vol. 83, 1986, Ikuta, T. et al "Three human alcohol dehydrogenase subunits: cDNA structure and molecular and evolutionary divergence" p. 634-638	1-6 (1864)
X	Proc. Natl. Acad. Sci. U.S.A., Vol. 85, 1988, Fukumoto, H. et al "Sequence, tissue distribution, and chromosomal localization of mRNA encoding a human glucose transporter-like protein" p. 5434-5438	1-6 (1878)

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X	J. Clin. Invest., Vol. 76, 1985, Cooke, N.E. et al. "Serum vitamin D-binding protein is a third member of the albumin and alpha fetoprotein gene family" p. 2420-2424	1-6 (1888)
X	J. Biol. Chem., Vol. 264, 1989, Huang, S.-H. et al. "Human deoxycytidine kinase: Sequence of cDNA clones and analysis of expression in cell lines with and without enzyme activity" p. 14762-14768	1-6 (1894)
X	J. Biol. Chem., Vol. 266, 1991, Huang, S.-H. et al. "Additions and corrections Human deoxycytidine kinase. Sequence of cDNA clones and analysis of expression in cell lines with and without anzyme activity" p. 5353-5353	1-6 (1894)
X	Somat. Cell Mol. Genet., Vol. 11, 1985, Bell, G.I. et al. "Human alpha-2-macroglobulin gene is located on chromosome 12" p. 285-289	1-6 (1895)
X	Proc. Natl. Acad. Sci. U.S.A., Vol. 81, 1984, Yang, F. et al. "Human transferrin: cDNA characterization and chromosomal localization" p. 2752-2756	1-6 (1902)
X	Proc. Natl. Acad. Sci. U.S.A., Vol. 83, 1986, Ny, T. et al. "Cloning and sequence of a cDNA coding for the human beta-migrating endothelial-cell-type plasminogen activator inhibitor" p. 6776-6780	1-6 (1904)
X	J. Biol. Chem., Vol. 267, 1992, Bausch-Jurken, M. T. et al "Molecular cloning of AMP deaminase isoform L: Sequence and bacterial expression of human AMPD2 cDNA" p. 22407-22413	1-6 (1908)
X	Gene, Vol. 44, 1986, Board, P. G. et al. "Molecular cloning and nucleotide sequence of human alpha-1 acid glycoprotein cDNA" p. 127-131	1-6 (1921)
X	Eur. J. Biochem., Vol. 155, 1986, Wathelet, M. et al. "Molecular cloning, full-length sequence and preliminary characterization of a 56-kDa protein induced by human interferons" p. 11-17	1-6 (2101)

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International application No.

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X	Biochemistry, Vol. 25, 1986, Koide, T. et al. "Amino acid sequence of human histidine-rich glycoprotein derived from the nucleotide sequence of its cDNA" p. 2220-2225	1-6 (2174)
X	Biochemistry, Vol. 22, 1983, Friezner-Degen, S. J. et al. "Characterization of the complementary deoxyribonucleic acid and gene coding for human prothrombin" p. 2087-2097	1-6 (2214)
X	Biochem. J., Vol. 268, 1990, Steinkasserer, A. et al. "Heterogeneity in human serum amyloid A protein. Five different variants from one individual demonstrated by cDNA sequence analysis." p. 287-193	1-6 (2238)
X	Nucleic Acids Res., Vol. 17, 1989, Fabrizi, G. M. et al. "Sequence of a cDNA specifying subunit VIIa of human cytochrome c oxidase" p. 7107-7107	1-6 (2264)
X	Proc. Natl. Acad. Sci. U.S.A., Vol. 86, 1989, Sims, J. E. et al. "Cloning of the interleukin 1 receptor from human T cells" p. 8946-8950	1-6 (2265)
X	Eur. J. Biochem., Vol. 169, 1987, Mackinnon, C. M. et al. "Molecular cloning of cDNA for human complement component Cls. The complete amino acid sequence" p. 547-553	1-6 (2266)
X	J. Virol., Vol. 65, 1990, Tsujimoto, A. et al. "Isolation of cDNA for DNA binding proteins which specifically bind to TAX-responsive enhancer element in the LTR of HTLA-1" p. 1420-1426	1-6 (2475)
X	Immunogenetics, Vol. 37, 1993, Emi, N. et al. "Isolation of a novel cDNA clone showing marked similarity to ME491/CD63 superfamily" p. 193-198	1-6 (2556)
X	Nature, Vol. 353, 1991, Kelly, A. P. et al. "A new human HLA class II-related locus, DM" p. 571-573	1-6 (2583)

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International application No.
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Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	J. Biol. Chem., Vol. 265, 1990, Hla, T. et al. "An abundant transcript induced in differentiating human endothelial cells encodes a polypeptide with structural similarities to G-protein-coupled receptors" p. 9308-9313	1-6 (2600)
X	J. Biol. Chem., Vol. 267, 1992, White, R. T. et al. "Human adipsin is identical to complement factor D and expressed at high levels in adipose tissue" p. 9210-9213	1-6 (2802)
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X	Nucleic Acids Res., Vol. 17, 1989, Sawada, R. et al. "Complementary DNA sequence and deduced peptide sequence for CD59/MEM43 antigen, the human homologue of murine lymphocyte antigen Ly-6c" p. 6728-6728	1-6 (2954)
X	DNA Cell Biol., Vol. 9, 1990, Sawada, R. et al. "Isolation and expression of the full-length cDNA encoding CD59 antigen of human lymphocytes" p. 213-220	1-6 (2954)
X	Proc. Natl. Acad. Sci. U.S.A., Vol. 87, 1990, Weller, P. A. et al. "Complete sequence of human vinculin and assignment of the gene to chromosome 10" p. 5667-5671	1-6 (2983)
X	Cell, Vol. 58, 1989, Mellentin, J. D. et al. "LYL-1, a novel gene involved by chromosomal translocation in T-cell leukemia, codes for a protein with a helix-loop-helix DNA binding motif" p. 77-83	1-6 (3023)
X	Cell, Vol. 60, 1990, Uze, G. et al. "Genetic transfer of a functional human interferon alpha receptor into mouse cells: Cloning and expression of its cDNA" p. 225-234	1-6 (3041)
X	Biochem. Biophys. Res. Commun., Vol. 179, 1991, Xiao, L. et al. "Characterization of a full length cDNA which codes for the human spermidine/spermine N-1-acetyltransferase" p. 407-415	1-6 (3053)

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International application No.
PCT/JP94/01916

C (Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	J. Biol. Chem., Vol. 266, 1991, Casero, R. A. Jr. et al. "Isolation and characterization of a cDNA clone that codes for human spermidine/spermine N-1-acetyltransferase" p. 810-814	1-6 (3053)
X	Nucleic Acids Res., Vol. 20, 1992, Wintzerith, M. et al. "Sequence of the human RNA polymerase II largest subunit" p. 910-910	1-6 (3083)
X	J. Cell Biol., Vol. 103, 1986, Lawler, J. et al. "The Structure of Human Thrombospondin, an adhesive Glycoprotein with Multiple Calcium binding Sites and Homologies with Several Different Proteins" p. 1635-1648	1-6 (3266)
X	Nature, Vol. 352, 1991, Maslen, C. L. et al. "Partial sequence of a candidate gene for the marfan syndrome" p. 334-337	1-6 (3334)
X	J. Cell Biol., Vol. 111, 1990, Fishman, G. I. et al. "Molecular Characterization and Functional Expression of the Human Cardiac Gap Junction Channel" p. 589-598	1-6 (3403)
X	Cell, Vol. 40, 1985, Ebina, Y. et al. "The human insulin receptor cDNA: The structural basis for hormone-activated membrane signalling" p. 747-758	1-6 (3447)
X	Oncogene, Vol. 5, 1990, Westin, E. H. et al. "Alternative splicing of the human c-myb gene" p. 1117-1124	1-6 (3529)
X	Genomics, Vol. 4, 1989, Todd, S. et al. "cDNA sequence, interspecies comparison and gene mapping analysis of argininosuccinate lyase" p. 53-59	1-6 (3575)
X	FEBS Lett., Vol. 207, 1986, Codina, J. et al. "-Subunits of the human liver Gs/Gi signal-transducing proteins and those of bovine retinal rod cell transducin are identical" p. 187-192	1-6 (3796)
X	Nucleic Acids Res., Vol. 18, 1990, Roessler, B. J. et al. "Cloning of two distinct copies of human phosphoribosyl pyrophosphate synthetase cDNA" p. 193-193	1-6 (3828)

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International application No.

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Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	J. Biochem., Vol. 109, 1991, Sonoda, T. et. al. "Complete nucleotide sequence of human phosphoribosyl pyrophosphate synthetase subunit I (PRS I) cDNA and a comparison with human and rat PRPS gene families" p. 361-364	1-6 (3828)
X	J. Biol. Chem., Vol. 263, 1988, Wermuth, B. et al. "Human carbonyl reductase: Nucleotide sequence analysis of a cDNA and amino acid sequence of the encoded protein" p. 16185-16188	1-6 (4033)
X	Biochim. Biophys. Acta. Vol. 1048, 1990, Forrest, G. L. et al. "Induction of a human carbonyl reductase gene located on chromosome 21" p. 149-155	1-6 (4033)
X	Proc. Natl. Acad. Sci. U.S.A., Vol. 88, 1991, Schuetz, T. J. et al. "Isolation of a cDNA for HSF2: Evidence for two heat shock factor genes in humans' p. 6911-6915	1-6 (4093)
X	Nucleic Acids Res., Vol. 13, 1985, Hallewell, R. A. et al. "Human Cu/Zn superoxide dismutase cDNA: isolation of clones synthesising high levels of active or inactive enzyme from an expression library" p. 2017-2034	1-6 (4110)
X	Proc. Natl. Acad. Sci. U.S.A., Vol. 80, 1983, Sherman, L. et al. "Nucleotide sequence and expression of human chromosome 21 - encoded superoxide dismutase mRNA" p. 5465-5469	1-6 (4110)
X	J. Biol. Chem., Vol. 268, 1993, David, V. et al. "Interaction with newly synthesized and retained proteins in the endoplasmic reticulum suggests a chaperone function for human integral membrane protein IP90 (calnexin)" p. 9585-9592	1-6 (4373)
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X	J. Biol. Chem., Vol. 263, 1988, Verma, A. K. et al. "Complete primary structure of a human plasma membrane Ca ²⁺ pump" p. 14152-14159	1-6 (4673)
X	J. Biol. Chem., Vol. 267, 1992, Shechter, I. et al. "Solubilization, purification and characterization of a truncated form of rat hepatic squalene synthetase" p. 8628-8635	1-6 (4818)
X	J. Biol. Chem., Vol. 267, 1992, Mckenzie, T. L. et al. "Molecular cloning, expression, and characterization of the cDNA for the rat hepataic squalene synthase" p. 21368-21374	1-6 (4818)
X	Nucleic Acids Res., Vol. 13, 1985, Furutani, Y. et al. "Cloning and characterization of the cDNAs for human and rabbit interleukin-1 precursor" p. 5869-5882	1-6 (4872)
X	Proc. Natl. Acad. Sci U.S.A., Vol. 89, 1992, Katoh, M. et al. "K-sam gene encodes secreted as well as transmembrane receptor tyrosine kinase" p. 2960-2964	1-6 (4914)
X	Differentiation, Vol. 42, 1989, Kuruc, N. et al. "Synthesis of cytokeratin 13, a component characteristic of internal stratified epithelia, is not induced in human epidermal tumors" p. 111-123	1-6 (5264)
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X	J. Biol. Chem., Vol. 265, 1990, Opiplari, A. W. et al. "The A20 cDNA induced by tumor necrosis factor alpha-encodes a novel type of zinc finger protein" p. 14705-14708	1-6 (5427)
X	J. Biol. Chem., Vol. 265, 1990, McLean, J. W. et al. "cDNA sequence of the human integrin beta-5 subunit" p. 17126-17131	1-6 (5715)
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X	J. Biol. Chem., Vol. 261, 1986, Romeo, P. -H. et al. "Molecular cloning and nucleotide sequence of a complete human uroporphyrinogen decarboxylase cDNA" p. 9825-9831	1-6 (6569)
X	J. Cell Biol., Vol. 106, 1988, Leube, R. E. et al. "Molecular characterization and expression of the stratification-related cytokeratins 4 and 15" p. 1249-1261	1-6 (6875)
X	Proc. Natl. Acad. Sci. U.S.A., Vol. 85, 1988, Daher, K. A. et al. "Isolation and characterization of human defensin cDNA clones" p. 7327-7331	1-6 (7106)
X	J. Exp. Med., Vol. 172, 1990, Larsen, A. et al. "Expression Cloning of a Human Granulocyte Colony-stimulating Factor Receptor: a Structural Mosaic of Hematopoietin Receptor, Immunoglobulin, and Fibronectin Domains" p. 1559-1570	1-6 (7126)

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X	Oncogene, Vol. 8, 1993, Schulz, A. S. et al. "The genomic structure of the human UFO receptor" p. 509-513	1-6 (7790)
A	Nature Genetics, Vol. 2, 1992, Okubo, K. et al. "Large scale cDNA sequencing for analysis of quantitative and qualitative aspects of gene expression" p. 173-179	1-6
A	Nature Genetics, Vol. 2, 1992, Khan, A. S. et al. "Single pass sequencing and physical and genetic mapping of human brain cDNAs" p. 180-188	1-6

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